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(54) Title: CELL CYCLE PROGRESSION PROTEINS

(57) Abstract: Polynucleotides encoding a number of *Drosophila* gene products are provided. Polynucleotide probes derived from these nucleotide sequences, polypeptides encoded by the polynucleotides and antibodies that bind to the polypeptides are also provided.



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CELL CYCLE PROGRESSION PROTEINS

The present invention relates to a number of genes implicated in the processes of cell cycle progression, including mitosis and meiosis.

We have now identified a large number of genes in *Drosophila*, mutations in
5 which disrupt cell cycle progression, for example the processes of mitosis and/or meiosis. We have determined the phenotypes of these mutations and recovered nucleotide sequences associated with the corresponding genes. Many of these nucleotide sequences correspond to protein open reading frames (ORFs) present in the *Drosophila* genome.

Accordingly the present invention provides in one aspect a polynucleotide selected
10 from: (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 1 to 70 or the complement thereof; (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 1 to 70, or a fragment thereof; (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 1 to 70 or
15 a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

There is provided, according to another aspect of the present invention, a polynucleotide selected from: (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 1 to 14 or the complement thereof; (b) polynucleotides
20 comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 1 to 14, or a fragment thereof; (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 1 to 14 or a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined
25 in (a), (b) or (c).

We provide, according to yet a further aspect of the present invention, a polynucleotide selected from: (a) polynucleotides comprising any one of the nucleotide

sequences set out in Examples 15 to 19 or the complement thereof; (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 15 to 19, or a fragment thereof; (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in
5 Examples 15 to 19 or a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

As a further aspect of the present invention, there is provided a polynucleotide selected from: (a) polynucleotides comprising any one of the nucleotide sequences set out
10 in Examples 20 to 30 or the complement thereof; (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 20 to 30, or a fragment thereof; (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 20 to 30 or a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence
15 which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

We provide, according to a yet further aspect of the present invention, a polynucleotide selected from: (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 31 to 53 or the complement thereof; (b) polynucleotides
20 comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 31 to 53, or a fragment thereof; (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in 31 to 53 or a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a),
25 (b) or (c).

The present invention, in a further aspect, provides a polynucleotide selected from: (a) polynucleotides comprising any one of the nucleotide sequences set out in 54 to 70 or the complement thereof; (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in 54 to 70, or a fragment thereof; (c)

polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in 54 to 70 or a fragment thereof; (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

- 5 A polynucleotide probe which comprises a fragment of at least 15 nucleotides of a polynucleotide according to any of the above aspects of the invention.

 The present invention also provides a polypeptide which comprises any one of the amino acid sequences set out in Examples 1 to 70 or in any of Examples 1 to 14, Examples 15 to 19, Examples 20 to 30, Examples 31 to 53 and Examples 54 to 70, or a
10 homologue, variant, derivative or fragment thereof.

 Preferably the polypeptide is encoded by a cDNA sequence obtainable from a eukaryotic cDNA library, preferably a metazoan cDNA library (such as insect or mammalian) said DNA sequence comprising a DNA sequence being selectively detectable with a *Drosophila* nucleotide sequence as shown in any one of Examples 1 to 70.

- 15 The term "selectively detectable" means that the cDNA used as a probe is used under conditions where a target cDNA of the invention is found to hybridize to the probe at a level significantly above background. The background hybridization may occur because of other cDNAs present in the cDNA library. In this event background implies a level of signal generated by interaction between the probe and a non-specific cDNA
20 member of the library which is less than 10 fold, preferably less than 100 fold as intense as the specific interaction observed with the target cDNA. The intensity of interaction may be measured, for example, by radiolabelling the probe, e.g. with ³²P. Suitable conditions may be found by reference to the Examples, as well as in the detailed description below.

 A polynucleotide encoding a polypeptide of the invention is also provided.

- 25 The present invention further provides a vector comprising a polynucleotide of the invention, for example an expression vector comprising a polynucleotide of the invention

operably linked to a regulatory sequence capable of directing expression of said polynucleotide in a host cell.

Also provided is an antibody capable of binding a polypeptide of the invention.

5 In a further aspect the present invention provides a method for detecting the presence or absence of a polynucleotide of the invention in a biological sample which method comprises: (a) bringing the biological sample containing DNA or RNA into contact with a probe comprising a nucleotide of the invention under hybridising conditions; and (b) detecting any duplex formed between the probe and nucleic acid in the sample.

10 In another aspect the invention provides a method for detecting a polypeptide of the invention present in a biological sample which comprises: (a) providing an antibody of the invention; (b) incubating a biological sample with said antibody under conditions which allow for the formation of an antibody-antigen complex; and (c) determining whether antibody-antigen complex comprising said antibody is formed.

15 Knowledge of the genes involved in cell cycle progression allows the development of therapeutic agents for the treatment of medical conditions associated with aberrant cell cycle progression. Accordingly, the present invention provides a polynucleotide of the invention for use in therapy. The present invention also provides a polypeptide of the invention for use in therapy. The present invention further provides an antibody of the
20 invention for use in therapy.

In a specific embodiment, the present invention provides a method of treating a tumour or a patient suffering from a proliferative disease, comprising administering to a patient in need of treatment an effective amount of a polynucleotide, polypeptide and/or antibody of the invention.

25 The present invention also provides the use of a polypeptide of the invention in a method of identifying a substance capable of affecting the function of the corresponding

gene. For example, in one embodiment the present invention provides the use of a polypeptide of the invention in an assay for identifying a substance capable of inhibiting cell cycle progression. The substance may inhibit any of the steps or stages in the cell cycle, for example, formation of the nuclear envelope, exit from the quiescent phase of the cell cycle (G0), G1 progression, chromosome decondensation, nuclear envelope breakdown, START, initiation of DNA replication, progression of DNA replication, termination of DNA replication, centrosome duplication, G2 progression, activation of mitotic or meiotic functions, chromosome condensation, centrosome separation, microtubule nucleation, spindle formation and function, interactions with microtubule motor proteins, chromatid separation and segregation, inactivation of mitotic functions, formation of contractile ring, and cytokinesis functions. For example, possible functions of genes of the invention for which it may be desired to identify substances which affect such functions include chromatin binding, formation of replication complexes, replication licensing, phosphorylation or other secondary modification activity, proteolytic degradation, microtubule binding, actin binding, septin binding, microtubule organising centre nucleation activity and binding to components of cell cycle signalling pathways.

In a further aspect the present invention provides a method for identifying a substance capable of binding to a polypeptide of the invention, which method comprises incubating the polypeptide with a candidate substance under suitable conditions and determining whether the substance binds to the polypeptide.

In an additional aspect, the invention provides kits comprising polynucleotides, polypeptides or antibodies of the invention and methods of using such kits in diagnosing the presence of absence of polynucleotides and polypeptides of the invention including deleterious mutant forms.

Also provided is a substance identified by the above methods of the invention. Such substances may be used in a method of therapy, such as in a method of affecting cell cycle progression, for example mitosis and/or meiosis.

The invention also provides a process comprising the steps of: (a) performing one of the above methods; and (b) preparing a quantity of those one or more substances identified as being capable of binding to a polypeptide of the invention.

Also provided is a process comprising the steps of: (a) performing one of the above
5 methods; and (b) preparing a pharmaceutical composition comprising one or more substances identified as being capable of binding to a polypeptide of the invention.

We further provide a method for identifying a substance capable of modulating the function of a polypeptide of the invention or a polypeptide encoded by a polynucleotide of the invention, the method comprising the steps of: incubating the polypeptide with a
10 candidate substance and determining whether activity of the polypeptide is thereby modulated.

A substance identified by a method or assay according to any of the above methods or processes is also provided, as is the use of such a substance in a method of inhibiting the function of a polypeptide. Use of such a substance in a method of regulating a cell
15 division cycle function is also provided.

DETAILED DESCRIPTION OF THE INVENTION

The practice of the present invention will employ, unless otherwise indicated, conventional techniques of chemistry, molecular biology, microbiology, recombinant DNA and immunology, which are within the capabilities of a person of ordinary skill in
20 the art. Such techniques are explained in the literature. See, for example, J. Sambrook, E. F. Fritsch, and T. Maniatis, 1989, *Molecular Cloning: A Laboratory Manual*, Second Edition, Books 1-3, Cold Spring Harbor Laboratory Press; Ausubel, F. M. et al. (1995 and periodic supplements; *Current Protocols in Molecular Biology*, ch. 9, 13, and 16, John Wiley & Sons, New York, N.Y.); B. Roe, J. Crabtree, and A. Kahn, 1996, *DNA Isolation
25 and Sequencing: Essential Techniques*, John Wiley & Sons; J. M. Polak and James O'D. McGee, 1990, *In Situ Hybridization: Principles and Practice*; Oxford University Press; M. J. Gait (Editor), 1984, *Oligonucleotide Synthesis: A Practical Approach*, Irl Press; and, D.

M. J. Lilley and J. E. Dahlberg, 1992, *Methods of Enzymology: DNA Structure Part A: Synthesis and Physical Analysis of DNA* Methods in Enzymology, Academic Press. Each of these general texts is herein incorporated by reference.

Preferably, the polypeptides and polynucleotides of the invention are such that they
5 give rise to or are associated with defined phenotypes when mutated.

For example, mutations in the polypeptides and polynucleotides of the invention may be associated with a failure to complete cytokinesis; such polypeptides and polynucleotides are conveniently categorised as "Category 1". Phenotypes associated with Category 1 polypeptides and polynucleotides include any one or more of the following,
10 singly or in combination: Mitotic defects in brain: cytokinesis defect (polyploidy); Male semi-sterile, Meiotic defects in testis: cytokinesis defects, segregation defects.(Seg-01/62); Meiotic defects in testis: cytokinesis defects, abnormal spindles.(Ab-02/12); Mitotic defects in brain: cytokinesis defect (no overcondensation of diploids, high polyploidy); Meiotic defects in testis: cytokinesis defects. Dark bands in eyes, dominant; Meiotic
15 defects in testis: cytokinesis defects; Meiotic defects in testis:segregation defect, cytokinesis defect(Ck-09/32); Mitotic defects in brain: cytokinesis defect (no overcondensation of diploids, very high polyploidy); Mitotic defects in brain: cytokinesis defect(very high polyploidy); Mitotic defects in brain: cytokinesis defect. Meiotic defects in testis: cytokinesis defects (Mitotic higher level of condensation, polyploidy, Meiotic: Ck05/07); Mitotic defects in brain, Cytokinesis defect (no overcondensation of diploids, high polyploidy); Mitotic defects in brain: cytokinesis defect (very high polyploidy, chromosomes entangled?); Mitotic defects in brain: cytokinesis defect (very high polyploidy; Meiotic defects in testis: cytokinesis defects (Ck-04/06) `; Female sterile (anaphase bridges, lagging chromosomes); Mitotic defects in brain: cytokinesis defect.
20 Meiotic defects in testis: cytokinesis defects:(mitotic: high polyploidy, no diploids, higher mitotic index, meiotic: Ck-01/05); Meiotic defects in testis: cytokinesis defects; Meiotic defects in testis: cytokinesis defects(Ck-06/09); Meiotic defects in testis: segregation defects, cytokinesis defect(Ck-07/35); Meiotic defects in testis: cytokinesis defects.

Alternatively, mutations in the polypeptides and polynucleotides of the invention may be associated with a failure to enter M-phase; such polypeptides and polynucleotides are conveniently categorised as "Category 2". Phenotypes associated with Category 2 polypeptides and polynucleotides include any one or more of the following, singly or in combination: Meiotic defects in testis: no division(no meiosis); Mitotic defects in brain: no mitosis; Meiotic defects in testis: segregation defects, meiotic failure(Mf-07/75); Meiotic defects in testis: segregation defects, meiotic failure(Mf-05/31); Meiotic defects in testis: cytokinesis defects, meiotic failure(Mf-02/15).

Mutations in the polypeptides and polynucleotides of the invention may be associated with a metaphase arrest phenotype ("Category 3"). Phenotypes associated with Category 3 polypeptides and polynucleotides include any one or more of the following, singly or in combination: Mitotic defects in brain: prometaphase arrest (overcondensation, polyploidy, scattered chromosomes with bipolar spindle); Male sterile, Female sterile, Mitotic defects in brain: prometaphase arrest (Overcondensation, polyploidy, fewer anaphases, high mitotic index, scattered chromosomes with bipolar spindle); Mitotic defects in brain: (weak overcondensation, metaphase with bipolar spindle); Mitotic defects in brain: prometaphase arrest; Mitotic defects in brain: metaphase arrest; Mitotic defects in brain: metaphase arrest. (overcondensation, polyploidy, aneuploidy, few anaphases, high mitotic index, metaphase with bent bipolar spindle); Mitotic defects in brain: metaphase arrest.(overcondensation, polyploidy, few anaphases, high mitotic index, metaphase with bent bipolar spindle); Mitotic defects in brain: Metaphase arrest (overcondensation, polyploidy, aneuploidy, no anaphases, high mitotic index, metaphase with bipolar spindle); Mitotic defects in brain: metaphase arrest (overcondensation, metaphase with bipolar spindle; Meiotic defects in testis: segregation defects, multipolar spindles (Mul-02/29); Meiotic defects in testis: cytokinesis defects, abnormal spindles.(Ab-01/03); Mitotic defects in brain: metaphase arrest; Mitotic defects in brain: metaphase arrest (overcondensation, polyploidy, metaphase with bipolar spindle); Mitotic defects in brain: metaphase arrest. Meiotic defects in testis: segregation defects. Abnormal spindles (mitotic: High mitotic index, meiotic: Ab-08/24); Mitotic defects in brain: metaphase arrest(overcondensation, few anaphases, some polyploids); Mitotic defects in brain: prometaphase arrest (overcondensation, fewer anaphases, metaphase with bipolar spindle);

Mitotic defects in brain: metaphase arrest(condensation, no polyploidy, no anaphases, metaphase with bipolar spindle).

Mutations in Category 4 polypeptides and polynucleotides of the invention may be associated with an anaphase defect phenotype; phenotypes associated with Category 4 polypeptides and polynucleotides include any one or more of the following, singly or in combination: Mitotic defects in brain: anaphase defects (overcondensation, high polyploidy, some lagging chromosomes); Meiotic defects in testis: segregation defects; Male and female sterile, small wings, meiotic defects in testis: segregation defects, elongation defect; Mitotic defects in brain: anaphase defects(overcondensation, anaphase bridge, metaphase with swollen chromosomes and bipolar spindle); Mitotic defects in brain: Anaphase defects. (overcondensation, aneuploidy, some lagging chromosomes and breaks); Meiotic defects in testis: segregation defects; Meiotic defects in testis: segregation defects, multi-stage defects (PI-02/17); Meiotic defects in testis: segregation defects, multi-stage defects (PI-02/18); Meiotic defects in testis: cytokinesis defects, segregation defects (seg-01/01); Mitotic defects in brain: cytokinesis defect. Meiotic defects in testis: cytokinesis defect. Multi-stage defects Polyploidy, no overcondensation PI-01/10; Meiotic defects in testis: segregation defects, abnormal spindles. (Ab-03/30); Mitotic defects in brain: anaphase defects (weak, higher condensation, some polyploidy, fewer anaphases, polyploids with monopolar spindles); Mitotic defects in brain: anaphase defects (overcondensation, polyploidy (with overcondensation), few anaphases, metaphase with bipolar spindle); Meiotic defects in testis: cytokinesis defects; Meiotic defects in testis: segregation defects, multipolar spindles(Mul-02/22); Meiotic defects in testis: segregation defects, abnormal spindles (Ab-04/26); Meiotic defects in testis: cytokinesis defects, abnormal spindles (Ab-16/13); Mitotic defects in brain: anaphase defects. Meiotic defects in testis: segregation defects, abnormal spindles (mitotic : Overcondensation, lagging chromosomes/less aligned metaphase with bipolar spindles, Meiotic: Ab-06/20); Meiotic defects in testis: segregation defects; Meiotic defects in testis: no division (no meiosis); Meiotic defects in testis: segregation defects, abnormal spindles (Ab-12/48); Meiotic defects in testis: segregation defects, multipolar spindles(mitotic: High polyploids, no diploids, higher mitotic index Meiotic: Mul-02/59); Meiotic defects in testis: segregation defect; Meiotic defects in testis: segregation defects, abnormal spindles

(meiotic: Ab-08/42); Female sterile. Meiotic defects in testis: cytokinesis defects, segregation defects (Mitotic: Less condensed chromosomes, nuclear bridges, Meiotic: Seg-01/02; Mitotic defects in brain: anaphase defects; Meiotic defects in testis: cytokinesis defects, abnormal spindles (Ab-01/04); Meiotic defects in testis: segregation defects (overcondensation, fewer anaphases); Mitotic defects in brain: (some overcondensation, anaphase bridge, metaphase with swollen chromosome and bipolar spindle).

A fifth category ("Category 5") of polypeptides and polynucleotides of the invention are associated with the presence of small imaginal discs (block to proliferation). Phenotypes associated with Category 5 polypeptides and polynucleotides include any one or more of the following, singly or in combination: 2nd chromosome, small imaginal discs.

The polypeptides and polynucleotides of the invention may also be categorised according to their function, or their putative function.

For example, the polypeptides described here preferably comprise, and the polynucleotides described here are ones which preferably encode polypeptides comprising, any one or more of the following: a CBP activator protein; a CCR4-associated regulator of polymerase II transcription; a CTP synthase (CTPS); a Cyclin specific ubiquitin conjugating enzyme; a DNA packaging protein; a DNA repair protein; a DNA-binding protein involved in chromosomal organisation; a DNase IV; a EIF4G2 translation initiation factor; a eukaryotic translation initiation factor 6; a Ecdysone-induced protein 78C; a Egf2 translation factor; a G protein-coupled receptor kinase 7; a GTPase exchange factor; a phosphatidylinositol transfer protein beta isoform; a His-rich protein; a Lk6 kinase; a MAP kinase; a MAP kinase interacting kinase 1; a N-arginine dibasic convertase; a Phosphatidylinositol transfer protein; a RIP protein kinase; a RNA binding motif, single stranded interacting protein; a RNA binding protein; a RYK receptor tyrosine kinase; a Ribosomal protein L1; a selenide, water dikinase 1; a selenium donor protein 1; a selenophosphate synthetase 1; a Sqv-7-like protein; a sugar modification protein; a protein involved in cytokinesis and signalling; a TEK tyrosine kinase; a Translation elongation

factor; a UDP-galactose transporter; a v-erba related protein; a WD40 protein; a brahma protein; a calcium binding protein; a cell adhesion protein; a chaperone; a chromodomain helicase DNA binding protein; a chromodomain-helicase-DNA-binding protein; a coiled coil protein with ubiquitin like domain; a component of the 19S proteasome regulatory particle; a couch potato RNA binding protein; a cytidine 5-prime triphosphate synthetase; a cytoskeletal structural protein; a death domain containing protein; a developmentally expressed in axons of the CNS; a diacylglycerol-activated/phospholipid dependent protein kinase C inhibitor; a diazepam binding inhibitor; a diphosphate kinase; a dodecasattelite DNA binding protein; a doughnut protein tyrosine kinase; an elongation factor 2; a endoplasmic reticulum ATPase; a eukaryotic translation initiation factor 4E binding protein 2; a factor involved in axon guidance; a fatty-acid-Coenzyme A ligase; a flap structure-specific endonuclease 1; a protein involved in the formation of the contractile ring and the initiation of cytokinesis; a glucose-6-phosphate transporter; a glycoprotein glucosyltransferase; a growth factor; a transmembrane receptor protein tyrosine kinase involved in cell growth and maintenance; a guanyl-nucleotide exchange factor involved in signal transduction; a heat shock protein; a helicase; a high density lipoprotein binding protein; a histone acetyl transferase transcriptional activator; a histone acetyltransferase; a histone acetyltransferase GCN5; a protein involved in development of the abdomen (embryos); a protein involved in the development of the imaginal discs (larvae or pupae); a kinesin like protein 67a; a ligand-dependent nuclear receptor; a ligand-dependent nuclear receptor; a lola-like specific RNA polymerase II transcription factor; a matrix associated protein; a membrane glycoprotein; a mitotic heterochromatin fragment clone CH(2)6; a motor protein; a motor protein involved in cytoskeleton organization; a mushroom body RNA binding protein; a myosin like proteins; a nemo-like kinase; a non-ATPase protein; a nuclear receptor NR1E1; a nucleic acid binding protein; a nucleoside diphosphate kinase (NBR-A); a oly(rC)-binding protein 2 (hnRNP-E1); a peroxisome biogenesis factor 1; a phospholipid transporter involved in lipid metabolism; a phosphatase or enhancer of Pi uptake protein; a protease; a proteasome regulatory particle; a protein involved in cytoskeleton organization and/or biogenesis; a protein kinase associated with microtubules; a protein kinase mitogen-activated 7; a protein serine/threonine kinase involved in cell cycle, possibly targeted to cytoskeleton; a protein serine/threonine kinase involved in eye morphogenesis; a protein which associates with cdc25 phosphatase; a

protein which induces apoptosis; a ribonuclease P; a ribonuclease P protein subunit p29; a ser/thr phosphatase; a signal transduction protein; a signal transport protein; a sin3-associated polypeptide; a single stranded DNA/RNA binding protein; a sodium-dependent dicarboxylate transporters; a ssDNA/RNA binding proteins; a striatin, calmodulin-binding protein (STRN); a structural protein of ribosome involved in protein biosynthesis; a subtelomeric heterochromatin repeats; a sugar acetylase; a sugar modification protein; a suppressor of ras; a tRNA processing enzyme Ribonuclease P protein subunit; a thyroid hormone responsive gene; a tie receptor protein tyrosine kinase; a transacylase; a transcription factor; a transcription factor involved in chromatin remodelling; a transcriptional regulation of c-myc expression; a transcriptional regulator; a transcriptional regulators/telomeric silencing; a translation initiation factor; a tumor metastasis inhibitor; a tyrosine 3-monooxygenase/tryptophan 5-monooxygenase activation protein; a ubiquitin carrier protein; a ubiquitin-conjugating enzyme; a ugtUDP-glucose-glycoprotein glucosyltransferase; a zinc finger protein; an RNA polymerase II transcription factor; an acetylcholinesterase (YT blood group) precursor; an actin binding protein; an actin dependent regulator of chromatin; an acyl-CoA-binding protein; an alanine:glyoxylate aminotransferase; an alpha esterase; an ankyrin protein; an imitation-SWI protein; and an integrin beta 4 binding protein.

POLYPEPTIDES

It will be understood that polypeptides of the invention are not limited to polypeptides having the amino acid sequence set out in Examples 1 to 70 or fragments thereof but also include homologous sequences obtained from any source, for example related viral/bacterial proteins, cellular homologues and synthetic peptides, as well as variants or derivatives thereof.

Thus polypeptides of the invention also include those encoding homologues from other species including animals such as mammals (e.g. mice, rats or rabbits), especially primates, more especially humans. More specifically, homologues included within the scope of the invention include human homologues.

Thus, the present invention covers variants, homologues or derivatives of the amino acid sequence set out in Examples 1 to 70, as well as variants, homologues or derivatives of the nucleotide sequence coding for the amino acid sequences of the present invention.

5 In the context of the present invention, a homologous sequence is taken to include an amino acid sequence which is at least 15, 20, 25, 30, 40, 50, 60, 70, 80 or 90% identical, preferably at least 95 or 98% identical at the amino acid level over at least 50 or 100, preferably 200, 300, 400 or 500 amino acids with any one of the polypeptide sequences shown in the Examples. In particular, homology should typically be considered
10 with respect to those regions of the sequence known to be essential for protein function rather than non-essential neighbouring sequences. This is especially important when considering homologous sequences from distantly related organisms.

Although homology can also be considered in terms of similarity (i.e. amino acid residues having similar chemical properties/functions), in the context of the present
15 invention it is preferred to express homology in terms of sequence identity.

Homology comparisons can be conducted by eye, or more usually, with the aid of readily available sequence comparison programs. These publicly and commercially available computer programs can calculate % homology between two or more sequences.

% homology may be calculated over contiguous sequences, i.e. one sequence is
20 aligned with the other sequence and each amino acid in one sequence directly compared with the corresponding amino acid in the other sequence, one residue at a time. This is called an “ungapped” alignment. Typically, such ungapped alignments are performed only over a relatively short number of residues (for example less than 50 contiguous amino acids).

25 Although this is a very simple and consistent method, it fails to take into consideration that, for example, in an otherwise identical pair of sequences, one insertion or deletion will cause the following amino acid residues to be put out of alignment, thus

potentially resulting in a large reduction in % homology when a global alignment is performed. Consequently, most sequence comparison methods are designed to produce optimal alignments that take into consideration possible insertions and deletions without penalising unduly the overall homology score. This is achieved by inserting “gaps” in the
5 sequence alignment to try to maximise local homology.

However, these more complex methods assign “gap penalties” to each gap that occurs in the alignment so that, for the same number of identical amino acids, a sequence alignment with as few gaps as possible - reflecting higher relatedness between the two compared sequences - will achieve a higher score than one with many gaps. “Affine gap
10 costs” are typically used that charge a relatively high cost for the existence of a gap and a smaller penalty for each subsequent residue in the gap. This is the most commonly used gap scoring system. High gap penalties will of course produce optimised alignments with fewer gaps. Most alignment programs allow the gap penalties to be modified. However, it is preferred to use the default values when using such software for sequence comparisons.
15 For example when using the GCG Wisconsin Bestfit package (see below) the default gap penalty for amino acid sequences is -12 for a gap and -4 for each extension.

Calculation of maximum % homology therefore firstly requires the production of an optimal alignment, taking into consideration gap penalties. A suitable computer program for carrying out such an alignment is the GCG Wisconsin Bestfit package
20 (University of Wisconsin, U.S.A; Devereux *et al.*, 1984, Nucleic Acids Research 12:387). Examples of other software than can perform sequence comparisons include, but are not limited to, the BLAST package (see Ausubel *et al.*, 1999 *ibid* – Chapter 18), FASTA (Atschul *et al.*, 1990, J. Mol. Biol., 403-410) and the GENWORKS suite of comparison tools. Both BLAST and FASTA are available for offline and online searching (see
25 Ausubel *et al.*, 1999 *ibid*, pages 7-58 to 7-60). However it is preferred to use the GCG Bestfit program.

Although the final % homology can be measured in terms of identity, the alignment process itself is typically not based on an all-or-nothing pair comparison. Instead, a scaled similarity score matrix is generally used that assigns scores to each

pairwise comparison based on chemical similarity or evolutionary distance. An example of such a matrix commonly used is the BLOSUM62 matrix - the default matrix for the BLAST suite of programs. GCG Wisconsin programs generally use either the public default values or a custom symbol comparison table if supplied (see user manual for
5 further details). It is preferred to use the public default values for the GCG package, or in the case of other software, the default matrix, such as BLOSUM62.

Once the software has produced an optimal alignment, it is possible to calculate % homology, preferably % sequence identity. The software typically does this as part of the sequence comparison and generates a numerical result.

10 The terms “variant” or “derivative” in relation to the amino acid sequences of the present invention includes any substitution of, variation of, modification of, replacement of, deletion of or addition of one (or more) amino acids from or to the sequence providing the resultant amino acid sequence retains substantially the same activity as the unmodified sequence, preferably having at least the same activity as the polypeptides presented in the
15 sequence listings in the Examples.

Polypeptides having the amino acid sequence shown in the Examples, or fragments or homologues thereof may be modified for use in the present invention. Typically, modifications are made that maintain the biological activity of the sequence. Amino acid substitutions may be made, for example from 1, 2 or 3 to 10, 20 or 30 substitutions
20 provided that the modified sequence retains the biological activity of the unmodified sequence. Alternatively, modifications may be made to deliberately inactivate one or more functional domains of the polypeptides of the invention. Amino acid substitutions may include the use of non-naturally occurring analogues, for example to increase blood plasma half-life of a therapeutically administered polypeptide.

25 Conservative substitutions may be made, for example according to the Table below. Amino acids in the same block in the second column and preferably in the same line in the third column may be substituted for each other:

| | | |
|-----------|-------------------|---------|
| ALIPHATIC | Non-polar | G A P |
| | | I L V |
| | Polar - uncharged | C S T M |
| | | N Q |
| | Polar - charged | D E |
| | | K R |
| AROMATIC | | H F W Y |

Polypeptides of the invention also include fragments of the full length sequences mentioned above. Preferably said fragments comprise at least one epitope. Methods of identifying epitopes are well known in the art. Fragments will typically comprise at least 6 amino acids, more preferably at least 10, 20, 30, 50 or 100 amino acids.

5 Proteins of the invention are typically made by recombinant means, for example as described below. However they may also be made by synthetic means using techniques well known to skilled persons such as solid phase synthesis. Proteins of the invention may also be produced as fusion proteins, for example to aid in extraction and purification. Examples of fusion protein partners include glutathione-S-transferase (GST), 6xHis,
10 GAL4 (DNA binding and/or transcriptional activation domains) and β -galactosidase. It may also be convenient to include a proteolytic cleavage site between the fusion protein partner and the protein sequence of interest to allow removal of fusion protein sequences. Preferably the fusion protein will not hinder the function of the protein of interest sequence. Proteins of the invention may also be obtained by purification of cell extracts
15 from animal cells.

Proteins of the invention may be in a substantially isolated form. It will be understood that the protein may be mixed with carriers or diluents which will not interfere with the intended purpose of the protein and still be regarded as substantially isolated. A protein of the invention may also be in a substantially purified form, in which case it will
20 generally comprise the protein in a preparation in which more than 90%, e.g. 95%, 98% or 99% of the protein in the preparation is a protein of the invention.

A polypeptide of the invention may be labeled with a revealing label. The revealing label may be any suitable label which allows the polypeptide to be detected. Suitable labels include radioisotopes, e.g. ^{125}I , enzymes, antibodies, polynucleotides and linkers such as biotin. Labeled polypeptides of the invention may be used in diagnostic procedures such as immunoassays to determine the amount of a polypeptide of the invention in a sample. Polypeptides or labeled polypeptides of the invention may also be used in serological or cell-mediated immune assays for the detection of immune reactivity to said polypeptides in animals and humans using standard protocols.

A polypeptide or labeled polypeptide of the invention or fragment thereof may also be fixed to a solid phase, for example the surface of an immunoassay well or dipstick. Such labeled and/or immobilised polypeptides may be packaged into kits in a suitable container along with suitable reagents, controls, instructions and the like. Such polypeptides and kits may be used in methods of detection of antibodies to the polypeptides or their allelic or species variants by immunoassay.

Immunoassay methods are well known in the art and will generally comprise: (a) providing a polypeptide comprising an epitope bindable by an antibody against said protein; (b) incubating a biological sample with said polypeptide under conditions which allow for the formation of an antibody-antigen complex; and (c) determining whether antibody-antigen complex comprising said polypeptide is formed.

Polypeptides of the invention may be used in *in vitro* or *in vivo* cell culture systems to study the role of their corresponding genes and homologues thereof in cell function, including their function in disease. For example, truncated or modified polypeptides may be introduced into a cell to disrupt the normal functions which occur in the cell. The polypeptides of the invention may be introduced into the cell by *in situ* expression of the polypeptide from a recombinant expression vector (see below). The expression vector optionally carries an inducible promoter to control the expression of the polypeptide.

The use of appropriate host cells, such as insect cells or mammalian cells, is expected to provide for such post-translational modifications (e.g. myristolation,

glycosylation, truncation, lapidation and tyrosine, serine or threonine phosphorylation) as may be needed to confer optimal biological activity on recombinant expression products of the invention. Such cell culture systems in which polypeptides of the invention are expressed may be used in assay systems to identify candidate substances which interfere
5 with or enhance the functions of the polypeptides of the invention in the cell.

POLYNUCLEOTIDES

Polynucleotides of the invention include polynucleotides that comprise any one or more of the nucleic acid sequences set out in Examples 1 to 70 and fragments thereof. Polynucleotides of the invention also include polynucleotides encoding the polypeptides
10 of the invention. It will be understood by a skilled person that numerous different polynucleotides can encode the same polypeptide as a result of the degeneracy of the genetic code. In addition, it is to be understood that skilled persons may, using routine techniques, make nucleotide substitutions that do not affect the polypeptide sequence encoded by the polynucleotides of the invention to reflect the codon usage of any
15 particular host organism in which the polypeptides of the invention are to be expressed.

Polynucleotides of the invention may comprise DNA or RNA. They may be single-stranded or double-stranded. They may also be polynucleotides which include within them synthetic or modified nucleotides. A number of different types of modification to oligonucleotides are known in the art. These include methylphosphonate
20 and phosphorothioate backbones, addition of acridine or polylysine chains at the 3' and/or 5' ends of the molecule. For the purposes of the present invention, it is to be understood that the polynucleotides described herein may be modified by any method available in the art. Such modifications may be carried out in order to enhance the *in vivo* activity or life span of polynucleotides of the invention.

25 The terms "variant", "homologue" or "derivative" in relation to the nucleotide sequence of the present invention include any substitution of, variation of, modification of, replacement of, deletion of or addition of one (or more) nucleic acid from or to the

sequence. Preferably said variant, homologues or derivatives code for a polypeptide having biological activity.

As indicated above, with respect to sequence homology, preferably there is at least 50 or 75%, more preferably at least 85%, more preferably at least 90% homology to the sequences shown in the sequence listing herein. More preferably there is at least 95%, more preferably at least 98%, homology. Nucleotide homology comparisons may be conducted as described above. A preferred sequence comparison program is the GCG Wisconsin Bestfit program described above. The default scoring matrix has a match value of 10 for each identical nucleotide and -9 for each mismatch. The default gap creation penalty is -50 and the default gap extension penalty is -3 for each nucleotide.

The present invention also encompasses nucleotide sequences that are capable of hybridising selectively to the sequences presented herein, or any variant, fragment or derivative thereof, or to the complement of any of the above. Nucleotide sequences are preferably at least 15 nucleotides in length, more preferably at least 20, 30, 40 or 50 nucleotides in length.

The term "hybridization" as used herein shall include "the process by which a strand of nucleic acid joins with a complementary strand through base pairing" as well as the process of amplification as carried out in polymerase chain reaction technologies.

Polynucleotides of the invention capable of selectively hybridising to the nucleotide sequences presented herein, or to their complement, will be generally at least 70%, preferably at least 80 or 90% and more preferably at least 95% or 98% homologous to the corresponding nucleotide sequences presented herein over a region of at least 20, preferably at least 25 or 30, for instance at least 40, 60 or 100 or more contiguous nucleotides.

The term "selectively hybridizable" means that the polynucleotide used as a probe is used under conditions where a target polynucleotide of the invention is found to hybridize to the probe at a level significantly above background. The background

hybridization may occur because of other polynucleotides present, for example, in the cDNA or genomic DNA library being screening. In this event, background implies a level of signal generated by interaction between the probe and a non-specific DNA member of the library which is less than 10 fold, preferably less than 100 fold as intense as the
5 specific interaction observed with the target DNA. The intensity of interaction may be measured, for example, by radiolabelling the probe, e.g. with ^{32}P .

Hybridization conditions are based on the melting temperature (T_m) of the nucleic acid binding complex, as taught in Berger and Kimmel (1987, Guide to Molecular Cloning Techniques, Methods in Enzymology, Vol 152, Academic Press, San Diego CA), and
10 confer a defined "stringency" as explained below.

Maximum stringency typically occurs at about $T_m - 5^\circ\text{C}$ (5°C below the T_m of the probe); high stringency at about 5°C to 10°C below T_m ; intermediate stringency at about 10°C to 20°C below T_m ; and low stringency at about 20°C to 25°C below T_m . As will be understood by those of skill in the art, a maximum stringency hybridization can be used to
15 identify or detect identical polynucleotide sequences while an intermediate (or low) stringency hybridization can be used to identify or detect similar or related polynucleotide sequences.

In a preferred aspect, the present invention covers nucleotide sequences that can hybridise to the nucleotide sequence of the present invention under stringent conditions
20 (e.g. 65°C and $0.1\times\text{SSC}$ { $1\times\text{SSC} = 0.15\text{ M NaCl}$, $0.015\text{ M Na}_3\text{ Citrate pH } 7.0$ }).

Where the polynucleotide of the invention is double-stranded, both strands of the duplex, either individually or in combination, are encompassed by the present invention. Where the polynucleotide is single-stranded, it is to be understood that the complementary sequence of that polynucleotide is also included within the scope of the present invention.

25 Polynucleotides which are not 100% homologous to the sequences of the present invention but fall within the scope of the invention can be obtained in a number of ways. Other variants of the sequences described herein may be obtained for example by probing

DNA libraries made from a range of individuals, for example individuals from different populations. In addition, other viral/bacterial, or cellular homologues particularly cellular homologues found in mammalian cells (e.g. rat, mouse, bovine and primate cells), may be obtained and such homologues and fragments thereof in general will be capable of
5 selectively hybridising to the sequences shown in the Examples. Such sequences may be obtained by probing cDNA libraries made from or genomic DNA libraries from other animal species, and probing such libraries with probes comprising all or part of any one of the sequences shown in the Examples under conditions of medium to high stringency. The nucleotide sequences of the human homologues described in the Examples, may
10 preferably be used to identify other primate/mammalian homologues since nucleotide homology between human sequences and mammalian sequences is likely to be higher than is the case for the *Drosophila* sequences identified herein.

Similar considerations apply to obtaining species homologues and allelic variants of the polypeptide or nucleotide sequences of the invention.

15 Variants and strain/species homologues may also be obtained using degenerate PCR which will use primers designed to target sequences within the variants and homologues encoding conserved amino acid sequences within the sequences of the present invention. Conserved sequences can be predicted, for example, by aligning the amino acid sequences from several variants/homologues. Sequence alignments can be performed
20 using computer software known in the art. For example the GCG Wisconsin PileUp program is widely used.

The primers used in degenerate PCR will contain one or more degenerate positions and will be used at stringency conditions lower than those used for cloning sequences with single sequence primers against known sequences. It will be appreciated by the skilled
25 person that overall nucleotide homology between sequences from distantly related organisms is likely to be very low and thus in these situations degenerate PCR may be the method of choice rather than screening libraries with labeled fragments the sequences disclosed in the Examples.

In addition, homologous sequences may be identified by searching nucleotide and/or protein databases using search algorithms such as the BLAST suite of programs. This approach is described in the Examples.

Alternatively, such polynucleotides may be obtained by site directed mutagenesis of characterised sequences, such as the sequences disclosed in the Examples. This may be useful where for example silent codon changes are required to sequences to optimise codon preferences for a particular host cell in which the polynucleotide sequences are being expressed. Other sequence changes may be desired in order to introduce restriction enzyme recognition sites, or to alter the property or function of the polypeptides encoded by the polynucleotides. For example, further changes may be desirable to represent particular coding changes found in the sequences disclosed in the Examples which give rise to mutant genes which have lost their regulatory function. Probes based on such changes can be used as diagnostic probes to detect such mutants.

Polynucleotides of the invention may be used to produce a primer, e.g. a PCR primer, a primer for an alternative amplification reaction, a probe e.g. labeled with a revealing label by conventional means using radioactive or non-radioactive labels, or the polynucleotides may be cloned into vectors. Such primers, probes and other fragments will be at least 8, 9, 10, or 15, preferably at least 20, for example at least 25, 30 or 40 nucleotides in length, and are also encompassed by the term polynucleotides of the invention as used herein.

Polynucleotides such as a DNA polynucleotides and probes according to the invention may be produced recombinantly, synthetically, or by any means available to those of skill in the art. They may also be cloned by standard techniques.

In general, primers will be produced by synthetic means, involving a step wise manufacture of the desired nucleic acid sequence one nucleotide at a time. Techniques for accomplishing this using automated techniques are readily available in the art.

Longer polynucleotides will generally be produced using recombinant means, for example using a PCR (polymerase chain reaction) cloning techniques. This will involve making a pair of primers (e.g. of about 15 to 30 nucleotides) flanking a region of the lipid targeting sequence which it is desired to clone, bringing the primers into contact with mRNA or cDNA obtained from an animal or human cell, performing a polymerase chain reaction under conditions which bring about amplification of the desired region, isolating the amplified fragment (e.g. by purifying the reaction mixture on an agarose gel) and recovering the amplified DNA. The primers may be designed to contain suitable restriction enzyme recognition sites so that the amplified DNA can be cloned into a suitable cloning vector

Polynucleotides or primers of the invention may carry a revealing label. Suitable labels include radioisotopes such as ^{32}P or ^{35}S , enzyme labels, or other protein labels such as biotin. Such labels may be added to polynucleotides or primers of the invention and may be detected using by techniques known *per se*.

Polynucleotides or primers of the invention or fragments thereof labeled or unlabeled may be used by a person skilled in the art in nucleic acid-based tests for detecting or sequencing polynucleotides of the invention in the human or animal body.

Such tests for detecting generally comprise bringing a biological sample containing DNA or RNA into contact with a probe comprising a polynucleotide or primer of the invention under hybridising conditions and detecting any duplex formed between the probe and nucleic acid in the sample. Such detection may be achieved using techniques such as PCR or by immobilising the probe on a solid support, removing nucleic acid in the sample which is not hybridised to the probe, and then detecting nucleic acid which has hybridised to the probe. Alternatively, the sample nucleic acid may be immobilised on a solid support, and the amount of probe bound to such a support can be detected. Suitable assay methods of this and other formats can be found in for example WO89/03891 and WO90/13667.

Tests for sequencing nucleotides of the invention include bringing a biological sample containing target DNA or RNA into contact with a probe comprising a polynucleotide or primer of the invention under hybridising conditions and determining the sequence by, for example the Sanger dideoxy chain termination method (see
5 Sambrook *et al.*).

Such a method generally comprises elongating, in the presence of suitable reagents, the primer by synthesis of a strand complementary to the target DNA or RNA and selectively terminating the elongation reaction at one or more of an A, C, G or T/U residue; allowing strand elongation and termination reaction to occur; separating out
10 according to size the elongated products to determine the sequence of the nucleotides at which selective termination has occurred. Suitable reagents include a DNA polymerase enzyme, the deoxynucleotides dATP, dCTP, dGTP and dTTP, a buffer and ATP. Dideoxynucleotides are used for selective termination.

Tests for detecting or sequencing nucleotides of the invention in a biological
15 sample may be used to determine particular sequences within cells in individuals who have, or are suspected to have, an altered gene sequence, for example within cancer cells including leukaemia cells and solid tumours such as breast, ovary, lung, colon, pancreas, testes, liver, brain, muscle and bone tumours. Cells from patients suffering from a proliferative disease may also be tested in the same way.

20 In addition, the identification of the genes described in the Examples will allow the role of these genes in hereditary diseases to be investigated. In general, this will involve establishing the status of the gene (e.g. using PCR sequence analysis), in cells derived from animals or humans with, for example, neurological disorders or neoplasms.

The probes of the invention may conveniently be packaged in the form of a test kit
25 in a suitable container. In such kits the probe may be bound to a solid support where the assay format for which the kit is designed requires such binding. The kit may also contain suitable reagents for treating the sample to be probed, hybridising the probe to nucleic acid in the sample, control reagents, instructions, and the like.

NUCLEIC ACID VECTORS

Polynucleotides of the invention can be incorporated into a recombinant replicable vector. The vector may be used to replicate the nucleic acid in a compatible host cell. Thus in a further embodiment, the invention provides a method of making polynucleotides of the invention by introducing a polynucleotide of the invention into a replicable vector, introducing the vector into a compatible host cell, and growing the host cell under conditions which bring about replication of the vector. The vector may be recovered from the host cell. Suitable host cells include bacteria such as *E. coli*, yeast, mammalian cell lines and other eukaryotic cell lines, for example insect Sf9 cells.

Preferably, a polynucleotide of the invention in a vector is operably linked to a control sequence that is capable of providing for the expression of the coding sequence by the host cell, i.e. the vector is an expression vector. The term "operably linked" means that the components described are in a relationship permitting them to function in their intended manner. A regulatory sequence "operably linked" to a coding sequence is ligated in such a way that expression of the coding sequence is achieved under condition compatible with the control sequences.

The control sequences may be modified, for example by the addition of further transcriptional regulatory elements to make the level of transcription directed by the control sequences more responsive to transcriptional modulators.

Vectors of the invention may be transformed or transfected into a suitable host cell as described below to provide for expression of a protein of the invention. This process may comprise culturing a host cell transformed with an expression vector as described above under conditions to provide for expression by the vector of a coding sequence encoding the protein, and optionally recovering the expressed protein. Vectors will be chosen that are compatible with the host cell used.

The vectors may be for example, plasmid or virus vectors provided with an origin of replication, optionally a promoter for the expression of the said polynucleotide and

optionally a regulator of the promoter. The vectors may contain one or more selectable marker genes, for example an ampicillin resistance gene in the case of a bacterial plasmid or a neomycin resistance gene for a mammalian vector. Vectors may be used, for example, to transfect or transform a host cell.

5 Control sequences operably linked to sequences encoding the polypeptide of the invention include promoters/enhancers and other expression regulation signals. These control sequences may be selected to be compatible with the host cell for which the expression vector is designed to be used in. The term promoter is well-known in the art and encompasses nucleic acid regions ranging in size and complexity from minimal
10 promoters to promoters including upstream elements and enhancers.

The promoter is typically selected from promoters which are functional in mammalian cells, although prokaryotic promoters and promoters functional in other eukaryotic cells, such as insect cells, may be used. The promoter is typically derived from promoter sequences of viral or eukaryotic genes. For example, it may be a promoter
15 derived from the genome of a cell in which expression is to occur. With respect to eukaryotic promoters, they may be promoters that function in a ubiquitous manner (such as promoters of α -actin, β -actin, tubulin) or, alternatively, a tissue-specific manner (such as promoters of the genes for pyruvate kinase). They may also be promoters that respond to specific stimuli, for example promoters that bind steroid hormone receptors. Viral
20 promoters may also be used, for example the Moloney murine leukaemia virus long terminal repeat (MMLV LTR) promoter, the rous sarcoma virus (RSV) LTR promoter or the human cytomegalovirus (CMV) IE promoter.

It may also be advantageous for the promoters to be inducible so that the levels of expression of the heterologous gene can be regulated during the life-time of the cell.
25 Inducible means that the levels of expression obtained using the promoter can be regulated.

In addition, any of these promoters may be modified by the addition of further regulatory sequences, for example enhancer sequences. Chimeric promoters may also be

used comprising sequence elements from two or more different promoters described above.

Polynucleotides according to the invention may also be inserted into the vectors described above in an antisense orientation to provide for the production of antisense
5 RNA. Antisense RNA or other antisense polynucleotides may also be produced by synthetic means. Such antisense polynucleotides may be used in a method of controlling the levels of RNAs transcribed from genes comprising any one of the polynucleotides of the invention.

HOST CELLS

10 Vectors and polynucleotides of the invention may be introduced into host cells for the purpose of replicating the vectors/polynucleotides and/or expressing the polypeptides of the invention encoded by the polynucleotides of the invention. Although the polypeptides of the invention may be produced using prokaryotic cells as host cells, it is preferred to use eukaryotic cells, for example yeast, insect or mammalian cells, in
15 particular mammalian cells.

Vectors/polynucleotides of the invention may be introduced into suitable host cells using a variety of techniques known in the art, such as transfection, transformation and electroporation. Where vectors/polynucleotides of the invention are to be administered to animals, several techniques are known in the art, for example infection with recombinant
20 viral vectors such as retroviruses, herpes simplex viruses and adenoviruses, direct injection of nucleic acids and biolistic transformation.

PROTEIN EXPRESSION AND PURIFICATION

Host cells comprising polynucleotides of the invention may be used to express polypeptides of the invention. Host cells may be cultured under suitable conditions which
25 allow expression of the proteins of the invention. Expression of the polypeptides of the invention may be constitutive such that they are continually produced, or inducible, requiring a stimulus to initiate expression. In the case of inducible expression, protein

production can be initiated when required by, for example, addition of an inducer substance to the culture medium, for example dexamethasone or IPTG.

Polypeptides of the invention can be extracted from host cells by a variety of techniques known in the art, including enzymatic, chemical and/or osmotic lysis and
5 physical disruption.

Polypeptides of the invention may also be produced recombinantly in an *in vitro* cell-free system, such as the TnTTM (Promega) rabbit reticulocyte system.

ANTIBODIES

The invention also provides monoclonal or polyclonal antibodies to polypeptides of
10 the invention or fragments thereof. Thus, the present invention further provides a process for the production of monoclonal or polyclonal antibodies to polypeptides of the invention.

If polyclonal antibodies are desired, a selected mammal (e.g., mouse, rabbit, goat, horse, etc.) is immunised with an immunogenic polypeptide bearing an epitope(s) from a polypeptide of the invention. Serum from the immunised animal is collected and treated
15 according to known procedures. If serum containing polyclonal antibodies to an epitope from a polypeptide of the invention contains antibodies to other antigens, the polyclonal antibodies can be purified by immunoaffinity chromatography. Techniques for producing and processing polyclonal antisera are known in the art. In order that such antibodies may be made, the invention also provides polypeptides of the invention or fragments thereof
20 haptenised to another polypeptide for use as immunogens in animals or humans.

Monoclonal antibodies directed against epitopes in the polypeptides of the invention can also be readily produced by one skilled in the art. The general methodology for making monoclonal antibodies by hybridomas is well known. Immortal antibody-producing cell lines can be created by cell fusion, and also by other techniques such as
25 direct transformation of B lymphocytes with oncogenic DNA, or transfection with Epstein-Barr virus. Panels of monoclonal antibodies produced against epitopes in the

polypeptides of the invention can be screened for various properties; i.e., for isotype and epitope affinity.

An alternative technique involves screening phage display libraries where, for example the phage express scFv fragments on the surface of their coat with a large variety
5 of complementarity determining regions (CDRs). This technique is well known in the art.

Antibodies, both monoclonal and polyclonal, which are directed against epitopes from polypeptides of the invention are particularly useful in diagnosis, and those which are neutralising are useful in passive immunotherapy. Monoclonal antibodies, in particular, may be used to raise anti-idiotypic antibodies. Anti-idiotypic antibodies are
10 immunoglobulins which carry an "internal image" of the antigen of the agent against which protection is desired.

Techniques for raising anti-idiotypic antibodies are known in the art. These anti-idiotypic antibodies may also be useful in therapy.

For the purposes of this invention, the term "antibody", unless specified to the
15 contrary, includes fragments of whole antibodies which retain their binding activity for a target antigen. Such fragments include Fv, F(ab') and F(ab')₂ fragments, as well as single chain antibodies (scFv). Furthermore, the antibodies and fragments thereof may be humanised antibodies, for example as described in EP-A-239400.

Antibodies may be used in method of detecting polypeptides of the invention
20 present in biological samples by a method which comprises: (a) providing an antibody of the invention; (b) incubating a biological sample with said antibody under conditions which allow for the formation of an antibody-antigen complex; and (c) determining whether antibody-antigen complex comprising said antibody is formed.

Suitable samples include extracts tissues such as brain, breast, ovary, lung, colon,
25 pancreas, testes, liver, muscle and bone tissues or from neoplastic growths derived from such tissues.

Antibodies of the invention may be bound to a solid support and/or packaged into kits in a suitable container along with suitable reagents, controls, instructions and the like.

ASSAYS

The present invention provides assays that are suitable for identifying substances
5 which bind to polypeptides of the invention and which affect, for example, formation of the nuclear envelope, exit from the quiescent phase of the cell cycle (G0), G1 progression, chromosome decondensation, nuclear envelope breakdown, START, initiation of DNA replication, progression of DNA replication, termination of DNA replication, centrosome duplication, G2 progression, activation of mitotic or meiotic functions, chromosome
10 condensation, centrosome separation, microtubule nucleation, spindle formation and function, interactions with microtubule motor proteins, chromatid separation and segregation, inactivation of mitotic functions, formation of contractile ring, cytokinesis functions, chromatin binding, formation of replication complexes, replication licensing, phosphorylation or other secondary modification activity, proteolytic degradation,
15 microtubule binding, actin binding, septin binding, microtubule organising centre nucleation activity and binding to components of cell cycle signalling pathways.

In addition, assays suitable for identifying substances that interfere with binding of polypeptides of the invention, where appropriate, to components of cell division cycle machinery. This includes not only components such as microtubules but also signalling
20 components and regulatory components as indicated above. Such assays are typically *in vitro*. Assays are also provided that test the effects of candidate substances identified in preliminary *in vitro* assays on intact cells in whole cell assays. The assays described below, or any suitable assay as known in the art, may be used to identify these substances.

According to one aspect of the invention, therefore, we provide one or more
25 substances identified by any of the assays described below, *viz*, mitosis assays, meiotic assays, polypeptide binding assays, microtubule binding/polymerisation assays, microtubule purification and binding assays, microtubule organising centre (MTOC) nucleation activity assays, motor protein assay, assay for spindle assembly and function,

assays for dna replication, chromosome condensation assays, kinase assays, kinase inhibitor assays, and whole cell assays, each as described in further detail below.

CANDIDATE SUBSTANCES

A substance that inhibits cell cycle progression as a result of an interaction with a polypeptide of the invention may do so in several ways. For example, if the substance inhibits cell division, mitosis and/or meiosis, it may directly disrupt the binding of a polypeptide of the invention to a component of the spindle apparatus by, for example, binding to the polypeptide and masking or altering the site of interaction with the other component. A substance which inhibits DNA replication may do so by inhibiting the phosphorylation or de-phosphorylation of proteins involved in replication. For example, it is known that the kinase inhibitor 6-DMAP (6-dimethylaminopurine) prevents the initiation of replication (Blow, JJ, 1993, *J Cell Biol*122,993-1002). Candidate substances of this type may conveniently be preliminarily screened by *in vitro* binding assays as, for example, described below and then tested, for example in a whole cell assay as described below. Examples of candidate substances include antibodies which recognise a polypeptide of the invention.

A substance which can bind directly to a polypeptide of the invention may also inhibit its function in cell cycle progression by altering its subcellular localisation and hence its ability to interact with its normal substrate. The substance may alter the subcellular localisation of the polypeptide by directly binding to it, or by indirectly disrupting the interaction of the polypeptide with another component. For example, it is known that interaction between the p68 and p180 subunits of DNA polymerase alpha-primase enzyme is necessary in order for p180 to translocate into the nucleus (Mizuno et al (1998) *Mol Cell Biol*18,3552-62), and accordingly, a substance which disrupts the interaction between p68 and p180 will affect nuclear translocation and hence activity of the primase. A substance which affects mitosis may do so by preventing the polypeptide and components of the mitotic apparatus from coming into contact within the cell.

These substances may be tested using, for example the whole cells assays described below. Non-functional homologues of a polypeptide of the invention may also be tested

for inhibition of cell cycle progression since they may compete with the wild type protein for binding to components of the cell division cycle machinery whilst being incapable of the normal functions of the protein or block the function of the protein bound to the cell division cycle machinery. Such non-functional homologues may include naturally
5 occurring mutants and modified sequences or fragments thereof.

Alternatively, instead of preventing the association of the components directly, the substance may suppress the biologically available amount of a polypeptide of the invention. This may be by inhibiting expression of the component, for example at the level of transcription, transcript stability, translation or post-translational stability. An example
10 of such a substance would be antisense RNA or double-stranded interfering RNA sequences which suppresses the amount of mRNA biosynthesis.

Suitable candidate substances include peptides, especially of from about 5 to 30 or 10 to 25 amino acids in size, based on the sequence of the polypeptides described in the Examples, or variants of such peptides in which one or more residues have been
15 substituted. Peptides from panels of peptides comprising random sequences or sequences which have been varied consistently to provide a maximally diverse panel of peptides may be used.

Suitable candidate substances also include antibody products (for example, monoclonal and polyclonal antibodies, single chain antibodies, chimeric antibodies and
20 CDR-grafted antibodies) which are specific for a polypeptide of the invention. Furthermore, combinatorial libraries, peptide and peptide mimetics, defined chemical entities, oligonucleotides, and natural product libraries may be screened for activity as inhibitors of binding of a polypeptide of the invention to the cell division cycle machinery, for example mitotic/meiotic apparatus (such as microtubules). The candidate substances
25 may be used in an initial screen in batches of, for example 10 substances per reaction, and the substances of those batches which show inhibition tested individually. Candidate substances which show activity in *in vitro* screens such as those described below can then be tested in whole cell systems, such as mammalian cells which will be exposed to the inhibitor and tested for inhibition of any of the stages of the cell cycle.

Polypeptide Binding Assays

One type of assay for identifying substances that bind to a polypeptide of the invention involves contacting a polypeptide of the invention, which is immobilised on a solid support, with a non-immobilised candidate substance determining whether and/or to what extent the polypeptide of the invention and candidate substance bind to each other. Alternatively, the candidate substance may be immobilised and the polypeptide of the invention non-immobilised.

In a preferred assay method, the polypeptide of the invention is immobilised on beads such as agarose beads. Typically this is achieved by expressing the component as a GST-fusion protein in bacteria, yeast or higher eukaryotic cell lines and purifying the GST-fusion protein from crude cell extracts using glutathione-agarose beads (Smith and Johnson, 1988). As a control, binding of the candidate substance, which is not a GST-fusion protein, to the immobilised polypeptide of the invention is determined in the absence of the polypeptide of the invention. The binding of the candidate substance to the immobilised polypeptide of the invention is then determined. This type of assay is known in the art as a GST pulldown assay. Again, the candidate substance may be immobilised and the polypeptide of the invention non-immobilised.

It is also possible to perform this type of assay using different affinity purification systems for immobilising one of the components, for example Ni-NTA agarose and histidine-tagged components.

Binding of the polypeptide of the invention to the candidate substance may be determined by a variety of methods well-known in the art. For example, the non-immobilised component may be labeled (with for example, a radioactive label, an epitope tag or an enzyme-antibody conjugate). Alternatively, binding may be determined by immunological detection techniques. For example, the reaction mixture can be Western blotted and the blot probed with an antibody that detects the non-immobilised component. ELISA techniques may also be used.

Candidate substances are typically added to a final concentration of from 1 to 1000 nmol/ml, more preferably from 1 to 100 nmol/ml. In the case of antibodies, the final concentration used is typically from 100 to 500 µg/ml, more preferably from 200 to 300 µg/ml.

5 ***Microtubule Binding/Polymerisation Assays***

In the case of polypeptides of the invention that bind to microtubules, another type of *in vitro* assay involves determining whether a candidate substance modulates binding of a polypeptide of the invention to microtubules. Such an assay typically comprises contacting a polypeptide of the invention with microtubules in the presence or absence of
10 the candidate substance and determining if the candidate substance has an affect on the binding of the polypeptide of the invention to the microtubules. This assay can also be used in the absence of candidate substances to confirm that a polypeptide of the invention does indeed bind to microtubules. Microtubules may be prepared and assays conducted as follows:

15 ***Microtubule Purification and Binding Assays***

Microtubules are purified from 0-3h-old *Drosophila* embryos essentially as described previously (Saunders, *et al.*, 1997). About 3 ml of embryos are homogenized with a Dounce homogenizer in 2 volumes of ice-cold lysis buffer (0.1 M Pipes/NaOH, pH6.6, 5 mM EGTA, 1 mM MgSO₄, 0.9 M glycerol, 1 mM DTT, 1 mM PMSF, 1 µg/ml
20 aprotinin, 1 µg/ml leupeptin and 1 µg/ml pepstatin). The microtubules are depolymerized by incubation on ice for 15 min, and the extract is then centrifuged at 16,000 g for 30 min at 4°C. The supernatant is recentrifuged at 135,000 g for 90 min at 4°C. Microtubules in this later supernatant are polymerized by addition of GTP to 1 mM and taxol to 20 µM and incubation at room temperature for 30 min. A 3 ml aliquot of the extract is layered on top
25 of 3 ml 15% sucrose cushion prepared in lysis buffer. After centrifuging at 54,000g for 30 min at 20°C using a swing out rotor, the microtubule pellet is resuspended in lysis buffer.

Microtubule overlay assays are performed as previously described (Saunders *et al.*, 1997). 500 ng per lane of recombinant Asp, recombinant polypeptide, and bovine serum albumin (BSA, Sigma) are fractionated by 10% SDS-PAGE and blotted onto PVDF

membranes (Millipore). The membranes are preincubated in TBST (50mM Tris pH 7.5, 150 mM NaCl, 0.05% Tween 20) containing 5% low fat powdered milk (LFPM) for 1 h and then washed 3 times for 15 min in lysis buffer. The filters are then incubated for 30 minutes in lysis buffer containing either 1 mM GDP, 1 mM GTP, or 1 mM GTP- γ -S.

5 MAP-free bovine brain tubulin (Molecular Probes) is polymerised at a concentration of 2 μ g/ml in lysis buffer by addition of GTP to a final concentration of 1 mM and incubated at 37°C for 30 min. The nucleotide solutions are removed and the buffer containing polymerised microtubules added to the membranes for incubation for 1h at 37°C with addition of taxol at a final concentration of 10 μ M for the final 30 min. The blots are then
10 washed 3 times with TBST and the bound tubulin detected using standard Western blot procedures using anti- β -tubulin antibodies (Boehringer Mannheim) at 2.5 μ g/ml and the Super Signal detection system (Pierce).

It may be desirable in one embodiment of this type of assay to deplete the polypeptide of the invention from cell extracts used to produce polymerise microtubules.

15 This may, for example, be achieved by the use of suitable antibodies.

A simple extension to this type of assay would be to test the effects of purified polypeptide of the invention upon the ability of tubulin to polymerise *in vitro* (for example, as used by Andersen and Karsenti, 1997) in the presence or absence of a candidate substance (typically added at the concentrations described above). *Xenopus* cell-
20 free extracts may conveniently be used, for example as a source of tubulin.

Microtubule Organising Centre (MTOC) Nucleation Activity Assays

Candidate substances, for example those identified using the binding assays described above, may be screening using a microtubule organising centre nucleation activity assay to determine if they are capable of disrupting MTOCs as measured by, for
25 example, aster formation. This assay in its simplest form comprises adding the candidate substance to a cellular extract which in the absence of the candidate substance has microtubule organising centre nucleation activity resulting in formation of asters.

In a preferred embodiment, the assay system comprises (i) a polypeptide of the invention and (ii) components required for microtubule organising centre nucleation activity except for functional polypeptide of the invention, which is typically removed by immunodepletion (or by the use of extracts from mutant cells). The components themselves are typically in two parts such that microtubule nucleation does not occur until the two parts are mixed. The polypeptide of the invention may be present in one of the two parts initially or added subsequently prior to mixing of the two parts.

Subsequently, the polypeptide of the invention and candidate substance are added to the component mix and microtubule nucleation from centrosomes measured, for example by immunostaining for the polypeptide of the invention and visualising aster formation by immuno-fluorescence microscopy. The polypeptide of the invention may be preincubated with the candidate substance before addition to the component mix. Alternatively, both the polypeptide of the invention and the candidate substance may be added directly to the component mix, simultaneously or sequentially in either order.

The components required for microtubule organising centre formation typically include salt-stripped centrosomes prepared as described in Moritz *et al.*, 1998. Stripping centrosome preparations with 2 M KI removes the centrosome proteins CP60, CP190, CNN and γ -tubulin. Of these, neither CP60 nor CP190 appear to be required for microtubule nucleation. The other minimal components are typically provided as a depleted cellular extract, or conveniently, as a cellular extract from cells with a non-functional variant of a polypeptide of the invention. Typically, labeled tubulin (usually β -tubulin) is also added to assist in visualising aster formation.

Alternatively, partially purified centrosomes that have not been salt-stripped may be used as part of the components. In this case, only tubulin, preferably labeled tubulin is required to complete the component mix.

Candidate substances are typically added to a final concentration of from 1 to 1000 nmol/ml, more preferably from 1 to 100 nmol/ml. In the case of antibodies, the final

concentration used is typically from 100 to 500 µg/ml, more preferably from 200 to 300 µg/ml.

The degree of inhibition of aster formation by the candidate substance may be determined by measuring the number of normal asters per unit area for control untreated
5 cell preparation and measuring the number of normal asters per unit area for cells treated with the candidate substance and comparing the result. Typically, a candidate substance is considered to be capable of disrupting MTOC integrity if the treated cell preparations have less than 50%, preferably less than 40, 30, 20 or 10% of the number of asters found in untreated cells preparations. It may also be desirable to stain cells for γ -tubulin to
10 determine the maximum number of possible MTOCs present to allow normalisation between samples.

Motor Protein Assay

Polypeptides of the invention may interact with motor proteins such as the Eg5-like motor protein *in vitro*. The effects of candidate substances on such a process may
15 be determined using assays wherein the motor protein is immobilised on coverslips. Rhodamine labeled microtubules are then added and their translocation can be followed by fluorescent microscopy. The effect of candidate substances may thus be determined by comparing the extent and/or rate of translocation in the presence and absence of the candidate substance. Generally, candidate substances known to bind to a polypeptide of
20 the invention, would be tested in this assay. Alternatively, a high throughput assay may be used to identify modulators of motor proteins and the resulting identified substances tested for affects on a polypeptide of the invention as described above.

Typically this assay uses microtubules stabilised by taxol (e.g. Howard and Hyman 1993; Chandra and Endow, 1993 – both chapters in “Motility Assays for Motor Proteins”
25 Ed Jon Scholey, pub Academic Press). If however, a polypeptide of the invention were to promote stable polymerisation of microtubules (see above) then these microtubules could be used directly in motility assays.

Simple protein-protein binding assays as described above, using a motor protein and a polypeptide of the invention may also be used to confirm that the polypeptide of the invention binds to the motor protein, typically prior to testing the effect of candidate substances on that interaction.

5 ***Assay for Spindle Assembly and Function***

A further assay to investigate the function of polypeptide of the invention and the effect of candidate substances on those functions is an assay which measures spindle assembly and function. Typically, such assays are performed using *Xenopus* cell free systems, where two types of spindle assembly are possible. In the “half spindle” assembly
10 pathway, a cytoplasmic extract of CSF arrested oocytes is mixed with sperm chromatin. The half spindles that form subsequently fuse together. A more physiological method is to induce CSF arrested extracts to enter interphase by addition of calcium, whereupon the DNA replicates and kinetochores form. Addition of fresh CSF arrested extract then induces mitosis with centrosome duplication and spindle formation (for discussion of
15 these systems see Tournebize and Heald, 1996).

Again, generally, candidate substances known to bind to a polypeptide of the invention, or non-functional polypeptide variants of the invention, would be tested in this assay. Alternatively, a high throughput assay may be used to identify modulators of spindle formation and function and the resulting identified substances tested for affects
20 binding of the polypeptide of the invention as described above.

Assays for DNA Replication

Another assay to investigate the function of polypeptide of the invention and the effect of candidate substances on those functions is as assay for replication of DNA. A number of cell free systems have been developed to assay DNA replication. These can be
25 used to assay the ability of a substance to prevent or inhibit DNA replication, by conducting the assay in the presence of the substance. Suitable cell-free assay systems include, for example the SV-40 assay (Li and Kelly, 1984, *Proc. Natl. Acad. Sci USA* 81, 6973-6977; Waga and Stillman, 1994, *Nature* 369, 207-212.). A *Drosophila* cell free replication system, for example as described by Crevel and Cotteril (1991), *EMBO J.* 10,

4361-4369, may also be used. A preferred assay is a cell free assay derived from *Xenopus* egg low speed supernatant extracts described in Blow and Laskey (1986, *Cell* 47,577-587) and Sheehan et al. (1988, *J. Cell Biol.* 106, 1-12), which measures the incorporation of nucleotides into a substrate consisting of *Xenopus* sperm DNA or HeLa nuclei. The nucleotides may be radiolabelled and incorporation assayed by scintillation counting. Alternatively and preferably, bromo-deoxy-uridine (BrdU) is used as a nucleotide substitute and replication activity measured by density substitution. The latter assay is able to distinguish genuine replication initiation events from incorporation as a result of DNA repair. The human cell-free replication assay reported by Krude, et al (1997), *Cell* 88, 109-19 may also be used to assay the effects of substances on the polypeptides of the invention.

Other In Vitro Assays

Other assays for identifying substances that bind to a polypeptide of the invention are also provided. For example, substances which affect chromosome condensation may be assayed using the *in vitro* cell free system derived from *Xenopus* eggs, as known in the art.

Substances which affect kinase activity or proteolysis activity are of interest. It is known, for example, that temporal control of ubiquitin-proteasome mediated protein degradation is critical for normal G1 and S phase progression (reviewed in Krek 1998, *Curr Opin Genet Dev* 8, 36-42). A number of E3 ubiquitin protein ligases, designated SCFs (Skp1-cullin-F-box protein ligase complexes), confer substrate specificity on ubiquitination reactions, while protein kinases phosphorylate substrates destined for destruction and convert them into preferred targets for ubiquitin modification catalyzed by SCFs. Furthermore, ubiquitin-mediated proteolysis due to the anaphase-promoting complex/cyclosome (APC/C) is essential for separation of sister chromatids during mitosis, and exit from mitosis (Listovsky et al., 2000, *Exp Cell Res* 255, 184-191).

Substances which inhibit or affect kinase activity may be identified by means of a kinase assay as known in the art, for example, by measuring incorporation of ^{32}P into a suitable peptide or other substrate in the presence of the candidate substance. Similarly,

substances which inhibit or affect proteolytic activity may be assayed by detecting increased or decreased cleavage of suitable polypeptide substrates.

Assays for these and other protein or polypeptide activities are known to those skilled in the art, and may suitably be used to identify substances which bind to a polypeptide of the invention and affect its activity.

Whole Cell Assays

Candidate substances may also be tested on whole cells for their effect on cell cycle progression, including mitosis and/or meiosis. Preferably the candidate substances have been identified by the above-described *in vitro* methods. Alternatively, rapid throughput screens for substances capable of inhibiting cell division, typically mitosis, may be used as a preliminary screen and then used in the *in vitro* assay described above to confirm that the affect is on a particular polypeptide of the invention.

The candidate substance, i.e. the test compound, may be administered to the cell in several ways. For example, it may be added directly to the cell culture medium or injected into the cell. Alternatively, in the case of polypeptide candidate substances, the cell may be transfected with a nucleic acid construct which directs expression of the polypeptide in the cell. Preferably, the expression of the polypeptide is under the control of a regulatable promoter.

Typically, an assay to determine the effect of a candidate substance identified by the method of the invention on a particular stage of the cell division cycle comprises administering the candidate substance to a cell and determining whether the substance inhibits that stage of the cell division cycle. Techniques for measuring progress through the cell cycle in a cell population are well known in the art. The extent of progress through the cell cycle in treated cells is compared with the extent of progress through the cell cycle in an untreated control cell population to determine the degree of inhibition, if any. For example, an inhibitor of mitosis or meiosis may be assayed by measuring the proportion of cells in a population which are unable to undergo mitosis/meiosis and comparing this to the proportion of cells in an untreated population.

The concentration of candidate substances used will typically be such that the final concentration in the cells is similar to that described above for the *in vitro* assays.

A candidate substance is typically considered to be an inhibitor of a particular stage in the cell division cycle (for example, mitosis) if the proportion of cells undergoing that particular stage (i.e., mitosis) is reduced to below 50%, preferably below 40, 30, 20 or 10% of that observed in untreated control cell populations.

THERAPEUTIC USES

Many tumours are associated with defects in cell cycle progression, for example loss of normal cell cycle control. Tumour cells may therefore exhibit rapid and often aberrant mitosis. One therapeutic approach to treating cancer may therefore be to inhibit mitosis in rapidly dividing cells. Such an approach may also be used for therapy of any proliferative disease in general. Thus, since the polypeptides of the invention appear to be required for normal cell cycle progression, they represent targets for inhibition of their functions, particularly in tumour cells and other proliferative cells.

The term proliferative disorder is used herein in a broad sense to include any disorder that requires control of the cell cycle, for example, cardiovascular disorders such as restenosis and cardiomyopathy, auto-immune disorders such as glomerulonephritis and rheumatoid arthritis, dermatological disorders such as psoriasis, anti-inflammatory, anti-fungal, antiparasitic disorders such as malaria, emphysema and alopecia.

One possible approach is to express anti-sense constructs directed against polynucleotides of the invention, preferably selectively in tumour cells, to inhibit gene function and prevent the tumour cell from progressing through the cell cycle. Anti-sense constructs may also be used to inhibit gene function to prevent cell cycle progression in a proliferative cell. Another approach is to use non-functional variants of polypeptides of the invention that compete with the endogenous gene product for cellular components of cell cycle machinery, resulting in inhibition of function. Alternatively, compounds identified by the assays described above as binding to a polypeptide of the invention may

be administered to tumour or proliferative cells to prevent the function of that polypeptide. This may be performed, for example, by means of gene therapy or by direct administration of the compounds. Suitable antibodies of the invention may also be used as therapeutic agents.

5 Alternatively, double-stranded (ds) RNA is a powerful way of interfering with gene expression in a range of organisms that has recently been shown to be successful in mammals (Wianny and Zernicka-Goetz, 2000, Nat Cell Biol 2000, 2, 70-75). Double stranded RNA corresponding to the sequence of a polynucleotide according to the invention can be introduced into or expressed in oocytes and cells of a candidate organism
10 to interfere with cell division cycle progression.

 In addition, a number of the mutations described herein exhibit aberrant meiotic phenotypes. Aberrant meiosis is an important factor in infertility since mutations that affect only meiosis and not mitosis will lead to a viable organism but one that is unable to produce viable gametes and hence reproduce. Consequently, the elucidation of genes
15 involved in meiosis is an important step in diagnosing and preventing/treating fertility problems. Thus the polypeptides of the invention identified in mutant *Drosophila* having meiotic defects (as is clearly indicated in the Examples) may be used in methods of identifying substances that affect meiosis. In addition, these polypeptides, and corresponding polynucleotides, may be used to study meiosis and identify possible
20 mutations that are indicative of infertility. This will be of use in diagnosing infertility problems.

ADMINISTRATION

 Substances identified or identifiable by the assay methods of the invention may preferably be combined with various components to produce compositions of the
25 invention. Preferably the compositions are combined with a pharmaceutically acceptable carrier or diluent to produce a pharmaceutical composition (which may be for human or animal use). Suitable carriers and diluents include isotonic saline solutions, for example phosphate-buffered saline. The composition of the invention may be administered by

direct injection. The composition may be formulated for parenteral, intramuscular, intravenous, subcutaneous, intraocular or transdermal administration. Typically, each protein may be administered at a dose of from 0.01 to 30 mg/kg body weight, preferably from 0.1 to 10 mg/kg, more preferably from 0.1 to 1 mg/kg body weight.

5 Polynucleotides/vectors encoding polypeptide components (or antisense constructs) for use in inhibiting cell cycle progression, for example, inhibiting mitosis or meiosis, may be administered directly as a naked nucleic acid construct. They may further comprise flanking sequences homologous to the host cell genome. When the polynucleotides/vectors are administered as a naked nucleic acid, the amount of nucleic
10 acid administered may typically be in the range of from 1 µg to 10 mg, preferably from 100 µg to 1 mg. It is particularly preferred to use polynucleotides/ vectors that target specifically tumour or proliferative cells, for example by virtue of suitable regulatory constructs or by the use of targeted viral vectors.

Uptake of naked nucleic acid constructs by mammalian cells is enhanced by
15 several known transfection techniques for example those including the use of transfection agents. Example of these agents include cationic agents (for example calcium phosphate and DEAE-dextran) and lipofectants (for example lipofectamTM and transfectamTM). Typically, nucleic acid constructs are mixed with the transfection agent to produce a composition.

20 Preferably the polynucleotide, polypeptide, compound or vector described here may be conjugated, joined, linked, fused, or otherwise associated with a membrane translocation sequence.

Preferably, the polynucleotide, polypeptide, compound or vector, etc described here may be delivered into cells by being conjugated with, joined to, linked to, fused to, or
25 otherwise associated with a protein capable of crossing the plasma membrane and/or the nuclear membrane (i.e., a membrane translocation sequence). Preferably, the substance of interest is fused or conjugated to a domain or sequence from such a protein responsible for the translocational activity. Translocation domains and sequences for example include

domains and sequences from the HIV-1-trans-activating protein (Tat), *Drosophila* Antennapedia homeodomain protein and the herpes simplex-1 virus VP22 protein. In a highly preferred embodiment, the substance of interest is conjugated with penetratin protein or a fragment of this. Penetratin comprises the sequence

5 RQIKIWFQNRRMKWKK and is described in Derossi, *et al.*, (1994), *J. Biol. Chem.* 269, 10444-50; use of penetratin-drug conjugates for intracellular delivery is described in WO/00/01417. Truncated and modified forms of penetratin may also be used, as described in WO/00/29427.

10 Preferably the polynucleotide, polypeptide, compound or vector according to the invention is combined with a pharmaceutically acceptable carrier or diluent to produce a pharmaceutical composition. Suitable carriers and diluents include isotonic saline solutions, for example phosphate-buffered saline. The composition may be formulated for parenteral, intramuscular, intravenous, subcutaneous, intraocular or transdermal administration.

15 The routes of administration and dosages described are intended only as a guide since a skilled practitioner will be able to determine readily the optimum route of administration and dosage for any particular patient and condition.

The invention will now be further described by way of Examples, which are meant to serve to assist one of ordinary skill in the art in carrying out the invention and are not
20 intended in any way to limit the scope of the invention.

EXAMPLESGeneration and Identification of Lethal, Semi-Lethal and Sterile Third Chromosome Mutants Having Defects in Mitosis and/Or Meiosis, and Second Chromosome Mutants Having Defects in Imaginal Disc Development By P-Element5 Insertion Mutagenesis*P-element mutagenesis*

Transposable elements are widely used for mutagenesis in *Drosophila melanogaster* as they couple the advantages of providing effective genetic lesions with ease of detecting disrupted genes for the purpose of molecular cloning. To achieve near
 10 saturation of the genome with mutations resulting from mobilisation of the P-lacW transposon (a P-element marked with a mini-white gene, bearing the *E.coli lacZ* gene as an enhancer trap, and an *E.coli* replicon and ampicillin resistance gene to facilitate ‘plasmid rescue’ of sequences at the site of the P-insertion), *Drosophila* females that are homozygous for *P-lacW* (inserted on the X chromosome) are crossed with males carrying
 15 the transposase source P(Δ 2-3) (Deak et al., 1997). Random transpositions of the mutator element are then ‘captured’ in lines lacking transposase activity. Stable, or balanced, stocks bearing single lethal *P-lacW* insertions are made.

More than 41,000 lines are derived, of which approximately one-half are on the third chromosome. Originally some 3100 lethal or strong semi-lethal lines (in homozygous
 20 conditions) are identified. During preliminary characterisation unstable lines and clusters of the same mutation event are eliminated leaving 2460 lines to be characterised.

Screening for Mitotic and Meiotic Defects

About half of the mutants in the collection are embryonic lethals. We have carried out cytological screens of the 1155 lines that comprise late larval lethals, pupal lethals,
 25 pharate and adult semi-lethals for defective mitosis in the developing larval CNS. This has identified 69 mutations falling into 43 complementation groups that affect all stages of the mitotic cycle. The cytological screens involve examining orcein-stained squashed preparations of the larval CNS to detect abnormal mitotic cells. In lines where defects are

identified, the larval CNS is subjected to immunostaining to identify centromeres, spindle microtubules and DNA for further examination. This leads to clarification of the mitotic defect.

As a set of common functions are essential to both mitosis and meiosis, we then
5 identify mutations resulting in sterility and failed progression through male meiosis. This involves examining squashed preparations larval, pupal or adult testes by phase contrast microscopy. We examine “onion stage” spermatids in the 519 pupal and pharate lethal lines and 463 adult “semi-lethal” and viable lines for variations in size and number of nuclei which provides an indication of whether there have been defects in either
10 chromosome segregation or cytokinesis, respectively. A total of 54 lines of the 519 pupal and pharate lethal lines and 22 of the adult lines show such defects. However, another 67 lines show male sterility without having onion-stage defects. 12 lines showing onion stage defects have been scored as having mitotic defects in the independent cytological screen of squashed preparations of the larval CNS. Twelve further lines with onion stage defects
15 show female sterility and of these, 10 show maternal effect mitotic defects in syncytial embryos. Thus greater than one half of the meiotic mutants scored appear to represent cell division functions specific to male meiosis or have targeted male germ-line specific enhancer elements, thus revealing their meiotic function but in this test not their mitotic function.

20 Further characterisation of testis preparations of each line by phase-contrast microscopy with and without staining with Hoechst to reveal DNA defined 6 broad categories of meiotic mutants:

8 mutants from the collection show defects in meiotic entry or at early stages in the first meiotic division (MF1-8).

25 18 mutants (15 complementation groups) show abnormal meiotic spindles (AB1-16). Mutants in this group almost invariably show an associated weak defect in cytokinesis, and 7 show a strong defect in spermatid differentiation. 3 of these mutants

also show mitotic defects in larval brains or in embryos derived from homozygous mutant mothers.

18 mutants (16 complementation groups) also show abnormal meiotic spindles that are strongly multipolar (MUL1-15). Three of these also show maternal effect mitotic abnormalities of multipolar spindles in syncytial embryos.

4 mutants (3 complementation groups) show strong defects at all stages of spermatogenesis from the pre-meiotic stages to spermatid elongation stages (PL1-3). In this respect they resemble the *polo*¹ mutation.

4 mutants show segregation defects as indicated by spermatid nuclei of heterogeneous sizes (SEG1-4). The spindles appear normal but all have what are either chromosome bridges or lagging chromosomes. One of these also shows a maternal effect.

9 mutants (7 complementation groups) show predominant cytokinesis defects. Two complementation groups also have cytokinesis defects in mitotic cells in the larval brain.

In the Examples below, the designations MF, AB, MUL, PL, SEG or CK are included in the category description where available. Further phenotype information for each mutant described in the results section is provided in the “Phenotype” field. There is considerable overlap between these categories, and it will be of much interest to distinguish between mutants in which the primary defect results in secondary consequences, and mutants that affect more than one aspect of spermatogenesis, as for example appears to be the case with *polo* mutants (Sunkel and Glover, 1988; Carmena et al, 1998).

In the Examples, lines exhibiting mitotic and meiotic phenotypes are categorised generally into four categories:

Category 1 : Failure to complete cytokinesis

Category 2 : Failure to enter M-phase

Category 3: Metaphase arrest

Category 4: Anaphase defect

Category 5: Small Imaginal Discs (Block to Proliferation; see below)

5 Category 1 phenotypes are exhibited by mutations in Examples 1 to 14; while
Category 2 phenotypes are exhibited by mutations in Examples 15 to 19. Category 3
phenotypes are exhibited by mutations in Examples 20 to 30, Category 4 phenotypes are
exhibited by mutations in Examples 31 to 53. Mutations in Examples 54 to 74 exhibit a
Category 5 phenotype.

10 ***Generation and identification of second chromosome mutants having small or
no imaginal discs.***

In the case of the second chromosome the flies used were from a second
chromosome P-element collection established in Szeged, Hungary (Torok et al., 1993).
The process of P-element insertion mutagenesis is essentially as described above. 15475
15 insertions were recovered, of which 2711 were lethal or semi-lethal. After elimination of
clusters of identical mutants, 2399 lines representing 1748 independent lethal insertions
were recovered. Lines were chosen from the second chromosome collection on the basis
of having small or no imaginal discs, to indicate a disruption in cell cycle progression that
leads to underdevelopment of the discs. All the second chromosome mutants referred to in
20 the results section are noted under the "Phenotype" field as "second chromosome, small
imaginal discs" and comprise Category 5.

Cytological Mapping of the P-Element Insertion Sites

The site of insertion of the P-element in each mutant line was determined by *in situ*
hybridisation of P-element DNA to salivary gland polytene chromosomes as described in
25 Saunders et al., 1989. Wandering third stage larvae were dissected and fixed as described
and incubated with biotin-labeled DNA made from the *P-lacW* plasmid. After signal

detection chromosomes were stained with Giemsa and examined by microscopy and signals indicating the presence of P elements were assigned to polytene chromosome bands referring to the Bridges map (Lefevre, 1976). In the majority of cases a single P element was detected, only 10% of lines having multiple (two or three) insertions. The site of insertion is given as the "Map Position" field in the results section (for example 77B)

Plasmid Rescue of P-Elements from Mutant Drosophila Lines

Genomic DNA was isolated from adult flies by the method of Jowett et al., 1986, and plasmid rescue from the genomic DNA was performed according to Pirrotta et al., 1986. This allows the recovery of genomic DNA adjacent to the P-element which facilitates the identification of the site of P-element insertion and of genes which may be disrupted by the insertion. Essentially, genomic DNA derived from about 200 flies was digested with a restriction enzyme known to have a site within the P-element (EcoRI or SacII for cloning sequences to the left of the element, or XbaI, BglII, PstI or BamHI for sequences to the right of the element). The digested DNA was ligated overnight, and plasmids recovered by electroporation of the ligated DNA into *E.coli* XL1-blue competent cells. Appropriate primers from within the P-lacW sequence were used to determine the sequence of the genomic DNA flanking the element (on average, 400 bp of sequence were obtained). The rescue sequences are provided in the results section under the heading "Rescue sequence". Where more than one sequence was recovered, the orientation of each sequence is also given.

Sequence Analysis of P Element Insertion Lines

Sequences flanking the insertion site of the P-element were derived by P element rescue as described above. In some cases sequence was obtained from only one side of the insertion, while in other cases sequences were obtained from both sides of the insertion.

As a first step, each P element rescue sequence was used to search a total database of *Drosophila melanogaster* sequences (database of the Berkley *Drosophila* Genome project) using the BLASTN program (which compares a nucleic acid sequence with a nucleic acid database, (Altschul and Lipman 1990)) with default parameters.

The search may identify a number of different types of match including *Drosophila* ESTs, known *Drosophila* genes and cloned genomic regions.

The ability to identify genes already known to be essential for cell cycle progression using this approach was confirmed, in this example, by the rescue sequence
5 obtained from line 1324/8 which mapped to the 77B locus which was used to search the database. A BLASTN search identified a number of matching *Drosophila* ESTs, a match with the known cell cycle regulatory gene *polo* and a cloned genomic region designated CSC: AC018188. These matches are recorded in the results sections under the field headings “*Drosophila* ESTs”, “*Drosophila* gene hit” and “Genomic hit, Accession No.”,
10 respectively. Any entries under “*Drosophila* gene hit” are further annotated with “(BLASTN with Rescue sequence)” to show that the match was obtained using the rescue sequence rather than a *Drosophila* EST or genomic clone ORF (see below). Accession numbers of ESTs, genes and genomic clones are provided where known. Genomic clones designations often include the Genbank designation as part of a longer designation.
15 However the Genbank designation is always the code beginning with “AC” and followed by six digits.

Where an EST was identified, this was subsequently used to search using the BLASTX program (default parameters) against databases of sequences from *Drosophila* and *Homo sapiens* (databases of the National centre for Biotechnology Information
20 (NCBI), National Library of Medicine, National Institute of Health, USA). In the case of line 1104/16, the search identified a known human gene, phosphatidylinositol transfer protein (accession no. P48739) implying a novel function for this protein in cytokinesis. Human Homologues identified as a result of a BLASTX search using a *Drosophila* EST are shown in the results section under the heading “Human homologues” and annotated
25 with “(BLASTX with EST)”. *Drosophila* genes identified as a result of a BLASTX search using a *Drosophila* EST are shown in the results section under the heading “*Drosophila* gene hit” and annotated with “(BLASTX with EST)”.

Where no *Drosophila* gene was identified using the initial BLASTN search but a matching genomic clone was found (a Bac or P1 clone often in excess of 100 kilobases), a

20 kilobase segment of this genomic sequence (10 kilobases either side flanking the site of the P-element insertion) was subjected to a number of analyses.

If the rescue sequence matched sequences that lie within a known gene present within the genomic clone then these are presented under the heading “*Drosophila* gene hit
5 (BLASTN with Rescue sequence”. The known gene sequence was then used in a BLASTX search of a human database (NCBI – see above) to identify any human homologues. These are shown in the “Human homologue” field and annotated with “(BLASTX with *Drosophila* gene)”.

If the rescue sequence does not match any sequences that lie with a known gene
10 within the genomic clone then the occurrence of ORFs within the 20 kilobase genomic segment was predicted using the Genscan programme (Burge and Karlin, 1997). Where the P-element was observed to be inserted into the coding region or within the 5’ untranslated region (which we defined as within 2 kilobases of the predicted start of the coding region) we assume the P element to be capable of disrupting the expression of the
15 predicted gene. Each predicted open reading frame (or predicted coding sequence) was then used to search *Drosophila* and human databases using the TBLASTN program (compares a protein query sequence against a nucleotide sequence database dynamically translated in all reading frames) and/or the TBLASTX program (compares a nucleotide query sequence dynamically translated in all reading frames against a nucleotide sequence
20 database dynamically translated in all reading frames) to determine whether the predicted open reading frame corresponded to a known gene. Typically, TBLASTX is only used when no matches are found using TBLASTN.

Where the TBLASTN search found a known *Drosophila* gene, then this is indicated in the results in the “*Drosophila* gene hit” field, annotated with “(TBLASTN
25 with predicted ORF)”. The *Drosophila* gene sequence was then typically used to search a human database (NCBI – see above) to identify any human homologues using BLASTX. These are shown in the “Human homologue” field and annotated with “(BLASTX with *Drosophila* gene)”.

Where the TBLASTN and/or TBLASTX search found a known human gene, then this is indicated in the results in the “Human homologue” field, annotated with “(TBLASTN (or TBLASTX) with predicted ORF)”.

5 If the TBLASTN and/or TBLASTX search found no *Drosophila* or human genes, then it was assumed that the original ORF corresponds to a novel gene. If the TBLASTN search found no *Drosophila* genes but identified a human homologue, then it was assumed that the original ORF corresponds to a novel *Drosophila* homologue of a known human gene.

10 *Additional Sequence Analysis using the Annotated D. melanogaster Sequence (GadFly).*

Rescue sequences were also used to search the fully annotated version of the *Drosophila* genome (GadFly; Adams, et al., 2000, Science 287, 2185-2195), using GlyBLAST at the Berkeley *Drosophila* Genome Projects web site to identify the genome segment (usually approximately 200-250 kb) containing the P-element insertion site. The
15 graphic representation of the genomic fragment available at GadFly allows the identification of all real and theoretical genes which flank the site of insertion. Candidate genes where the P-element is either inserted within the gene or close to the 5' end of the gene were identified. In GadFly, the *Drosophila* genes are given the designation CG (Complete gene) and usually details of human homologues are also given. In most cases,
20 this data confirms the data derived from the sequence analysis procedure described above, and in some cases new data is obtained. Where available both sets of data are included in the individual Examples described below. To identify further candidate human homologues, BLASTP (amino acid query sequence against amino acid database) searches with *Drosophila* sequences are used against the human genome project database and also
25 the Ensembl dataset. The Ensembl dataset comprises GeneWise gene predictions using a protein template where possible or Genscan followed by BLAST confirmation via protein, cDNA or EST hits. These are matched using WUBLASTP with default parameters (Altschul et al., 1990, *J Mol Biol* 215, 403-10). The results are filtered to contain only potential homologues. Only matches with the identity of more than 50% and length of
30 more than 50 amino acids are included.

Confirmation of Cell Cycle Involvement of Candidate Genes Using Double Stranded RNA Interference (RNAi)

P-elements usually insert into the region 5' to a *Drosophila* gene. This means that there is sometimes more than one candidate gene affected, as the P-element can insert into the 5' regions of two diverging genes (one on each DNA strand). In order to confirm which of the candidate genes is responsible for the cell cycle phenotype observed in the fly line, we use the technique of double stranded RNA interference to specifically knock out gene expression in *Drosophila* cells in tissue culture (Clemens, et al., 2000, *Proc. Natl. Acad. Sci. USA*, 6499-6503). The overall strategy is to prepare double stranded RNA (dsRNA) specific to each gene of interest and to transfect this into Schneider's *Drosophila* line 2 to inhibit the expression of the particular gene. The dsRNA is prepared from a double stranded, gene specific PCR product with a T7 RNA polymerase binding site at each end. The PCR primers consist of 25-30 bases of gene specific sequence fused to a T7 polymerase binding site (TAATACGACTCACTATAGGGACA), and are designed to amplify a DNA fragment of around 500bp. Although this is the optimal size, the sequences in fact range from 450 bp to 650 bp. Where possible, PCR amplification is performed using genomic DNA purified from Schneider's *Drosophila* line 2 as a template. This is only feasible where the gene has an exon of 450 bp or more. In instances where the gene possesses only short exons of less than 450 bp, primers are designed in different exons and PCR amplification is performed using cDNA derived from Schneider's *Drosophila* line 2 as a template.

A sample of PCR product is analysed by horizontal gel electrophoresis and the DNA purified using a Qiagen QiaQuick PCR purification kit. 1µg of DNA is used as the template in the preparation of gene specific single stranded RNA using the Ambion T7 Megascript kit. Single stranded RNA is produced from both strands of the template and is purified and immediately annealed by heating to 90 degrees C for 15 mins followed by gradual cooling to room temperature overnight. A sample of the dsRNA is analysed by horizontal gel electrophoresis.

3µg of dsRNA is transfected into Schneider's *Drosophila* line 2 using the transfection agent, Transfect (Gibco) and the cells incubated for 72 hours prior to fixation.

The DNA content of the cells is analysed by staining with propidium iodide and standard FACS analysis for DNA content. The cells in G1 and G2/S phases of the cell cycle are visualised as two separate population peaks in normal cycling S2 cells. In each experiment, Red Fluorescent Protein dsRNA is used as a negative control. In some cases
 5 the phenotype is confirmed by fixing cells on poly-lysine covered slides which are then stained for DNA using DAPI and for tubulin using an anti-tubulin antibody YL1/2 and appropriate fluorescent secondary antibody to visualise aberrant mitoses.

It should be noted that RNAi could not confirm phenotype in all cases. This is to be expected as the method relies on the ability of dsRNA to prevent new protein
 10 expression. Consequently, it is necessary that S2 cells express the specific cDNA of the gene in question, and also that the protein is turned over rapidly. It would therefore probably be difficult to sufficiently reduce levels of very stable proteins using this approach.

The layout of a typical entry in the results section is shown below. Not all fields
 15 present in the actual results section contain information for each individual *Drosophila* line described.

TYPICAL RESULTS LAYOUT

| | | |
|----|---------------------|---|
| 20 | Line ID | - <i>Drosophila</i> line designation |
| | Category | - Description of phenotype |
| | Reversion | - R = revertant, NR = non revertant, ? = not determined |
| | Map Position | - according to the Bridges map (Lefevre, 1976). |

| | |
|----|------------------------|
| 25 | Rescue ID |
| | Rescue Sequence |
| | [nucleotide sequence] |

Genomic hit, Accession No.

| | |
|----|--|
| 30 | Associated ORF |
| | GENSCAN_predicted_peptide [results of Genscan - amino acid sequence] |
| | GENSCAN_predicted_CDS [results of Genscan nucleotide sequence] |

| | |
|----|-----------------------------------|
| 35 | <i>Drosophila</i> Gene Hit |
| | (BLASTN with rescue sequence) |

(TBLASTN (or TBLASTX) with predicted ORF)
(BLASTX with EST)

Human Homologue

5 (BLASTX with *Drosophila* gene)
(TBLASTN (or TBLASTX) with predicted ORF)
(BLASTX with EST)
Drosophila EST

10 **Annotated *Drosophila* genome genomic segment**
Annotated *Drosophila* genome Complete gene candidate
Human homologue of Complete gene candidate

Putative function Derived from homologies or *Drosophila* experimental data
15 **Confirmation by RNAi** Description of Facs analysis DNA content profile

A specific example is as follows:

20 **Line ID** 1324/8
Category Mitotic defects in brain: metaphase arrest
 (overcondensation, some circular chromosomes, no anaphases,
 very high mitotic index, metaphase (or less aligned) with bipolar
25 **Reversion** R
Map Position 77B

Rescue ID B1E
30 **Rescue Sequence**
GTTTTGCCCATCGATTGCACGAAAACCAAGCACAAAGCGGAGAACGCGCCGA
AACCGTTCGATTTTTTAAATGCCAAAATGAATTGGACGTGAAGCGTCAGCTGA
ATTGGTGTGCCCGTTTCGGTGGCTATCGCACACTTTCTGGTATTTATCGCGGTA
TTTTGTTGAGTGTTGAACAACAAATTCTATGGCCGTTACCCTTTTGAATTTACT
35 TACTGGCGTTTACTCTGTTCGAATTGAGCGCAATATTTTTTCCTATTGCTCTGC
GCAACACTGTGTTTTAACCGCTATTTATTTGAAAATCTACAAAAACTAACCGTT
TACATTTTTGAAATTTCCAAAAGGGTTTTCCATAAATTGAGTTTTACTAAAACC
AGTCCAACGGTCCAACCTTTATATTGTTAGAAGCCCCTTTTCCTAATTGGAATTG
GCTTGCAAACGTTTTTCCTGAATTTAAAAATACTGCCACCCTTGTTAATTGCAGG
40 TTTTCCGAATCCCTGATTTGTTGTTTTAAAAAGAAAATTTATTAGAAACAGCTA
TCTCAACC

Genomic hit, Accession No. CSC:AC018188
***Drosophila* Gene Hit** Polo (X63361)
45 **Human Homologue** BLASTX PLK-1 (P53350)
***Drosophila* EST** several including LD11851 (AA392613) which match polo

Annotated *Drosophila* genome genomic segment AE003514

Annotated *Drosophila* genome Complete gene candidate CG12306
Human homolog of Complete gene candidate 1e-169 1709658 P53350
PLK1_HUMAN
SERINE/THREONINE-
5 PROTEIN KINASE PLK
(PLK-1)

Putative function Serine/threonine kinase known to be required for mitosis

10 **Confirmation by RNAi** Reduced G1 and G2/M peaks indicating fewer cycling cells, microscopy analysis of DNA and tubulin staining identified monopolar spindles characteristic of polo mutation in *Drosophila*.

CATEGORY 1: FAILURE TO COMPLETE CYTOKINESIS**Example 1 (Category 1)**

5 **Line ID** 1031/14
 Category Mitotic defects in brain: cytokinesis defect
 (polyplody)
 Reversion R
 Map Position 74B

10 **Rescue ID** 2A3B
 Rescue Sequence 1
 CCCCGGAACATATGTTTCAGTGTGGCCGCAGCAGAGTTGTCAAAACACGCTCCC
 CAATGAAATAACCTAAATGTGCCATCACTGTTACTTAACAGTTTCTGTTACTTT
 TCTAGCGGCATGTCAAAAAAACAAAAATATAGAAAATGCTAAATATATATTG
15 GACTAATGTGTTTAAATGTAACCTTACACTAGTAACAGATCCCCATTAATAAAA
 GCCAAACTCTAAAATTCTGCCACAAGTACTATTTCTCACGTAACACCTTACTA
 ACGGATTTACATGATATCTACGACAAGAACTGTTTGCTGATATAAAATTGC
 TATCACCGCTTTCCGTAAACACTTTTACACTGATGGATTACAAGTTCAATTAAT
 ACATCAACTTACCTTAACAATTTTAAGACAATAACTCCCACAATTTAATT
20 CAACCTACACCGCTTGATAATCAGCTGTTCTGTACAAAAACAATAACACTGT
 TAACAACAGCGCACAGTGGATAATACAGTCCTAAAGGCAATATACCCATTG
 GCATTTTT

Rescue ID 2A3S
25 **Rescue Sequence 2**
 TTCCGGGGGAGAATGGCTGCGATTTTCGCGTCGGTAAAAATAGCAAATACTCGTTA
 ATGTGCTGTGGGAACGCTTCCTCCCCGGCCCCAAAGTGGCCCCGAAGAAAGTGA
 GCAAATGTGCGCGCCGCAAGATAGTCGCCGCCGAACAAACGATAGTGACGAAA
 GTGATTTAATTCAACTACCAGCACTCCCGCAAATACGATGAGTATGTCGCGCGG
30 CGGCAACACAACCTCTGGACTTGCAGCCGCTCCTGGCGGAGAGCGATGTCGGAA
 ACAGGGAGCTGGAGGAGAAGATGGGCGGATCGGCGGATCGGTCATCGCTGCTC
 GATGGATCCGGTTCGAAGGAGCTGAGTCACCGGGAACGCGAGGACTCGGCGTT
 GTTCGTCAAGAAGATCGGGAGCGCCTTGTTCTATGGCTTGTCCTCCTTCATGATT
 ACGGTGGTAAACAAGACGGTGCTTACCTCCTACCACTTCCCCTCGTTCCTGTTCC
35 TCAGCCTCGGGCAACTTACTGCTAGCATTGTGGTCCTGGGCATGGGCAAAGCGC
 CTGAAAATGGTGAACCTTTTCCCCTTTTGCAGAGGAATACCTTCGCCAAGATCTTT
 CCGCTGCCACTGATATTTCTGGGAAACATGATGTTTGGACTGGGTGGCACAAAA
 ACCTTGAGTCTGCCCATGTTTCGCAGCCCTACGAC

40 **Genomic hit, Accession No.** AC019515

Associated ORF
 Genscan ORF1 predicted sequences:>15:31:57|GENSCAN_predicted_peptide_4|373_aa

MSMSRGGNTTLDLQPLLAESDVGNRELEEKMGGSADRSSLDDGSGSKELSHRER
EDSALFVKKIGSALFYGLSSFMITVVNKT VLT SYHFPSFLFLSLGQLTASIVVLGMG
KRLKLVNFPPLQRNTFAKIFPLPLIFLGNNMMFGLGGTKTSLPLMFALRRFSILMT
MLELKLGLRPSNAVQVSVYAMIGGALLAASDDL SFNM RGYIYVMITNAL TASN
5 GVVVKKKLDTSEIGKYGLMYYNLSL FMPALALNYVTGNLDQALNFEQWNSV
FVVQFLLSCVMGFILSYSTILCTQFNSALTTTIVGCLKNICVTYLG MFIGGDYVFSW
LNCIGINISVLASLLYTYVTFRRKRAPDKQDHL PSTRGENV

>15:31:57|GENSCAN_predicted_CDS_4|1122_bp

10 atgagtatgtcgcgcggcggaacacaactctggacttcagccgctcctggcggagagcgatgtcggaaacaggagctgga
ggagaagatgggcggatcggcgatcggatcgcctgcctgatccgggtcgaaggagctgagtcaccgggaacgcgag
gactcggcggttctcgtcaagaagatcgggagcgccttcttatggcttgctcctcctcatgattacgggtgtaacaagacgggtgc
ttacctctaccacttccccctcgttcctgttcctcagcctcgggcaacttactgctagcattgtggctcctgggcattgggcaagcgcct
gaaattggtgaacttccccctcgtcagaggaataccttcgcccaagatcttccgctgccactgatatttctgggaaacatgatgttg
15 gactgggtggcacaaaaaccttgagtctgccatgttcgcagccctacgacgcttctctatcctgatgaccatgctgctggagctca
agatcctgggactgcgaccttcgaatgcgggtcaggtcagcgtatacgaatgatcgggtggagcgcctgctggcgcctctgatga
tctgtcctcaacatgaggggctacatctatgtgatgatacgaacgccttgaccgcctcgaatggcgtatatgtgaagaaaaaactc
gacacctcggagatcggaaagtacggcctaattgtactacaactcgcctgtttatgtttctgcctgccctggccctcaactatgttacag
ggaatctagatcaggcgctgaactttgaacaatggaatgactcagtggttggtgcagttcctgctcagttgcgttatgggttcac
20 ctatcgtacagcaccatcctgtgcacgcaattcaactcggcgctgaccaccaccattgtgggatgcctgaaaaacatctgcgtaac
atatctgggcatgttcattggagggcactacgtctctcgtggctcaactgtattgggatcaacatcagcgtgctggctagtctgctct
acacgtacgtcacttttcggcggaagcgggctcccgataagcaggaccacttgcccagcaccgcggcgagaatgtctag

25 **Human Homologue** (TBLASTN with ORF1): KIAA0260 gene (D87449) and putative
Sqv-7-like protein (AJ005866)
Drosophila EST CK00510 (AA140776)

30 **Annotated Drosophila genome genomic segment** AE003524
Annotated Drosophila genome Complete gene candidate CG3874 – novel glucose-6-
phosphate transporter

35 **Human homologue of Complete gene candidate** EMBL:D87449 protein
KIAA0260_id:BAA13390
gi:166578 Similar to a
C.elegans protein encoded in
cosmid C52E12 (U50135) and
Ensembl predicted gene
ENSG00000024527
Clone:AL133320
40 Contig:AL133320.00001
8.10E-95

45 **Putative function** Sugar modification protein similar to proteins involved in
Drosophila cytokinesis and signalling

Confirmation by RNAi Marked increased G1 and S peak indicating mainly arrest in
G1

Example 2 (Category 1)

Line ID 1066/5
Category Male semi-sterile, Meiotic defects in testis: cytokinesis defects, segregation defects.
 5 (Seg-01/62)
Reversion ?
Map Position 89B
 10 **Rescue ID** F9E
Rescue Sequence
 GTATACCATTAGAGAATATGATGAAGAAGGACTGTAAGAAGATCCTTCAGTG
 AATTTGACTGCTGACGTCGATCGGAACCTTGCTGCGCTGACGTACAAAATCGCG
 AAGTGAATAAATAATATGGATGAGACCCTGTTTCGCCGACATATACAATAGTG
 15 CTCAAGACCTAATGGAATTATACGTTAATAACCAGCCACATTTCTTAGATATTT
 CTAATATGAGCCATCTGCTGCAGGTTCTTTCCAATATCTAATTCTAGATCTTCT
 TCGAATACGACCTTTTTGGCCATGAAACGATGATTTGCCACTTCATTCACAAG
 CATTAAATTTGTCATGATTCTCTTAAGCGTGCACCTTTATCTGAAAGTCTGAACAG
 CTGGCTGCGAAATGGATCCCCGGGATTGGAGATGGCAAGTAAATCTGTCCTCG
 20 CTACAAACAAGTGGGCACCACTGGGCATTCGGGGAATAGGGATATGGGTTGG
 GAATGGGGATATATTGTGGCATTGGCGAAAGGTCGCTATGC

Genomic hit, Accession No. CSC:AC019750

25 **Associated ORF**
 >16:04:57|GENSCAN_predicted_peptide_4|418_aa
 MKPIPNESKGTAAVGDATVVHDTVCTLFAVELDPYLRSSMGMRTTRRAQSGALLL
 QLLAVADGGFAAHICACKCRLRLPHVTCCCNRNPFKATAKAKGQAVSSTKPNQL
 CFHGCCGWIITTKGETFTENSPSIMSGFAWERHSLGECVVVAGTEQILLIGRTLIGR
 30 MSHTQTDSTSPFVVDCHSQLCGSKCKCICVSVGFCVRPSCQRFDMKIVWANLAM
 QKRFLLGAAIADMCCRNSVIWCKLQLDPVKPIDERADGSGLALVTKVCDNNNIV
 HYVVVAGVTGSQSRSLQPLRSGQNESTEQWPRTKGEGGFNNNSRNNKHSAPT
 QEQQELWQKQLLQDQRDDCHASGSFQSASFAETRSFTFDDTTAHSEFCFRTRAEK
 RRILVLLETSIKLKPDKYATSGHTRRCAIGLLHSII
 35 >16:04:57|GENSCAN_predicted_CDS_4|1257_bp
 atgaaacccattcccaacgaatccaagggaacccttgccgcagttggagatgctactgttggtcatgacgtgtgtactttgttgccg
 tagagcttgatccctatctcaggagcagcatgggaatgaggacgcgtagagctcaaagcggcgctctgttattacagctccttgccg
 gttgccgatggaggtttgtgctcatattgtgcctgcaagtgtcggttcgttgccacatgtcacatgttgtgcaaccggaatcct
 40 ttcaaggcaactgcaaaagcaaaaggtcaggcggtcagctccactaaaccaaaccagctttgcttcacggctgctgtggctggat
 aattactaccaaaggtgaaacgttcaccgaaaactcgcccagcatcatgagcgggtttgcgtgggagcggcatagccttggtgagt
 gcgtgggttggtggaacggaacaaatcctgctgattggcaggacattgattggccgcatgagccatactcaaactgattcgacc
 agcccccttggtgtcgaactgtgcggtccaagtgcgaatgtatctgtgtatctgtaggtttctgtgtgcgccgtct
 tgtcagcgtttgacatgaaaatagtttgggccaacttggtatgcaaaagcgatttctattaggagccgcatcgccgacatgtgct
 45 gccgaaatcggtgatttggtgcaactgcagctagatccagtcgaagccaattgacgaaagagccgacggcagcggtcttgcaact
 ggttaccaaagtatgcgataacaataacatcgtccactatgtggtcgttggtgggttacgggcagtcagtcacggtcacggctgc

aacccctccgctccggccaaaacgagtcacagaacaatggccaaggacgaaggggggggaggggggattcaataacaaca
gcaggaacaacaacattctgctcccacgcaagagcagcaggaactgtggcaaaaacagctgctgcaggatcaacgagacgat
tgtcatgccagtggaagcttccagtctgcgtcattcgcgagacgcgtagtttcacgttcgacgacacaaccgctcacagcgaattt
tgtttcggactagagctgagaaacggcgaattttggtgcttctggaaacatcgattaaactaaaacccgataagtatgcgacaagc
5 ggtcacactcggcgatgtgcgataggattgctgcattcgattatag

***Drosophila* Gene Hit** rescue sequence: mitotic heterochromatin fragment clone CH(2)6
(L36595) and subtelomeric heterochromatin repeats (L03284).

10 **Human Homologue** TBLASTN with ORF1: nebula (nla) (AF147700)
BLASTX with nebula: Down Syndrome candidate region 1-like
protein 2 (AF176117)

***Drosophila* EST** rescue sequence: CK01138 (AA141069)

15 **Annotated *Drosophila* genome genomic segment** AE003712
Annotated *Drosophila* genome Complete gene candidate CG6072 - nebula
CG6046 – sap18

20 **Human homologue of Complete gene candidate** CG6072- 8e-36 'ZAKI4 a thyroid
hormone responsive gene in human
skin fibroblasts' also known as
DOWN SYNDROME CANDIDATE
REGION 1-LIKE 1; DSCR1L1
EMBL:D83407
25 protein_id:BAA11911 gi:143504

30 CG6046- 3e-45 2108210 (U96915)
sin3 associated polypeptide p18
[Homo sapiens] and gi5032067
C7E479FFE9CA5774
|ref|NP_005861.1| sin3-associated
polypeptide, 18kD [Homo sapiens]
(1.90E-43)

35 **Putative function** Nebula unknown function, Sap18 transcription factor

Confirmation by RNAi Both show reduction in G1 and G2/S peaks indicating fewer
cycling cells

| | | |
|----|----------------------------------|---|
| | Line ID | 234/50 |
| | Category | Meiotic defects in testis: cytokinesis defects, abnormal spindles. (Ab-02/12) |
| | Reversion | R |
| 5 | Map Position | 89B |
| | Rescue ID | 2C5E |
| | Rescue Sequence | |
| 10 | | GTTTGACTGCTGACGTCGATCGGAACTTGCTGCGCTGACGTACAAAATCGCGA AGTGAATAAATAATATGGATGAGACTCCTGTTTCGCCGACATATAACAATAGTG CTCAAGACCCTAATGGAATTATACGTTAATAACCAGCCACATTTCTTAGATAT TTCTAATATGAGCCATCTGCTGCAGGTTCTTTCCAATATCTAATTCTAGATCTT CTTCGAATACGACCTTTTTTGGCCATGAAACGATGATTTGCCACTTCATTCACAA GCATTAATTTGTCATGATTCTCTTAAGCGTGCACTTTATCTGAAAGTCTGAACA 15 GCTGGCTGCGAAATGGATTCCCCGGATTGGAGATGGCAAGTAAATCTGTCCTC GCTACAAACAAGTGGGCACCACTGGGCATTCGGGGAATAGGGATATGGGTTG GAAA |
| 20 | Drosophila EST | rescue sequence: CK01138 (AA141069) |
| | All other entries as for 1066/5. | |

Example 3 (Category 1)

Line ID 1104/16
Category Mitotic defects in brain: cytokinesis defect
5 (no overcondensation of diploids, high polyploidy)
Reversion R
Map Position 92A

Rescue ID B5P

10 **Rescue Sequence 1**
CTCCGGACACGCAGTAGCTAAATAACAACTCATTACTAGTATATTACTGCCG
CCGATTTGCAAGCGCGTACCGATCCCGATACCAGGCCAATCGCACTCCCCAGT
TGTACGTCATCACTTAAGTAATAAATCAGCGGCAAATCGCATAAATTGCTATT
GATATTCCGCCCGCTGTGTGTGCGTGTGTATTTGCAAGAGAGTGTGTGTGTGT
15 GTGTGCATATGACTCGTGC GTTTAGCCGACAATTGGAGAAAAAGCATTACCAA
TCCCAATTGGCTAACTAACTAAAGTTGGCTTGGCCAAACATAAACAAAAAGT
GCGGGCGCAGCGATTTGGCAGCGAAACATATACACCAAAGCGCTATTGGCAG
ATATATATGTAGATTAAATATAGAAAGTGCGTGCGAAGGTTAAGAGTCGAGT
GCAAGTGCATTTATATTTGGAAATAATAAATGCTACAAT

20 **Rescue ID** B5E

Rescue Sequence 2
GTCCGGAGCGGAGCTAAAGTTCGATGTTTCGTGCAAAACACTTCGATTCCGATA
GGCGGATGCTATCGATTTTCGGCGATGCCCGTTGGTCACACTTGGTGGTGGGCG
25 CTGCCCCTCGCCGACTATCGATAGCACAAAGCGGGTTATTTAGGTGTGCGCAGC
TTGTAAGGGTGACTCATGCTGTTAAAATTATTATAAAAAAGTTAATGAATATAA
TATAGTTATAATAAAATTATATATAAATCTATAAGATCAAAGATCATCAGTTA
TCATTTATCATTTGATTATATGAAAAACAAGAACAGAAACAAGATTTAATAGG
TTTTTGAAATGTGAAAATGTGGGTTACCCCCAATTCTTATTCGAAATTAAATAA
30 CCTAAAGAACAGTTATACACAGATAGGTAATTTGCACATAAGCCAAATTTTGT
CTAGAATTCCGCGGAATTAATTCTTGAAGACGAAAGGGCCTCCGTGATACGCC
TATTTTATAGGTTAATGTCATGATAATAATGGTTTCTTA

Genomic hit, Accession No. AC006589

35 **Associated ORF**
Genscan: ORF1 predicted sequences
>/tmp/aaaaainga|GENSCAN_predicted_peptide_2|850_aa
MATRGANVIWFRHGLRLHDNPALLAALADKDQGIALIPVFIFDGESAGTKNVGY
NRMRFLLDSLQDIDDQLQAATDGRGRLLVFEGEPAYIFRRLHEQVRLHRICIEQDC
40 EPIWNERDESIRSLCRELNIDFVEKVSHTLWDPQLVIETNGGIPPLTYQMFLIRCTH
HNGDVNGDEDTGEGEGTGGRIDWAKEGACWRAGNSDEQECQACQSVSSVIMM
VLQYSNNPAHHCQLLECLMTLKHN VVKDILCVVAYGTAVSRTSAAKLLFYYWP
AFNANLFDRKVLLSKLTNDLVPFTCQREHCPNSGNAEAAKV CYDHSISIA YAPDC
PPPLYLCIECAN EIHREHGSLEFGDILHPMQQVSMVCENKNCRSNEKSAFSICFSTE
45 CASFNGNHPIRYCSQCHSNRHNSRRGGDHVVHRS LQPAWQMDPEMQMHMVESV
VSLLREAKPLNFEPGKESSSES SKKNGSGITADNISLEERQRLGRYGIWLLVGRCTP

TADTPVEVLGRILSMLFHWFHVTA YSYDGFISCLVPHPEYARVGGHWETLASRT
 SHLKEGLQRLICLVPEYVITSEIWDYVMPHWMEAITNDVAEKELNELKIVLSKILD
 PEMSPLGFDAKTMYNFVAIRFEKTTAKVQQQALHWLQILTKLEILIPLVQLFAMF
 GDGVRIMKYGIQHELMREKDAQS QSLAKAPKTPCKESKETKADMANPPRPPVVE
 5 DDSGNTSAISDDEAPTNRHTEFSTDAEHNLTCCILMLDILLKQMELQDVEQHMG
 HTSVCENVSRLIKCMVTAARVGLSSHVCALKVPIEDIIEEEKSSRKSPPESDKEKTR
 DRDVSLSMAPLPIPLGPLGGFADP

>/tmp/aaaaainga|GENSCAN_predicted_CDS_2|2553_bp

10 atggccacgcgaggggcgaatgtgatttggttcgccatggattgcgcctccatgataat
 cccgctctattggccgccctcgccgataaggatcagggatagccctaattcccgtttcatattcgatggagagagtgcaggtacc
 aagaatgtgggttacaatcgatgcgtttcctcctggactcgttgaggacatcgatgatcagctacaggcggcaactgatggacg
 tggacgcctcctggtcttcgagggcgaaccggcttatacttcgccggctacatgagcaagtgcgtctgcacaggatttgcatag
 agcaggactgcgagccaatttgaatgagcgcgatgaaagcatccgttctctatgtcgggagctgaatatcgactttgtcgagaag
 15 gtatcacacacgcttgggatccgcaattggtgattgagaccaatggtggcattccaccgctgacctaccaaatgttctgatacgt
 gcacgcaccacaatggagatgtgaatggggatgaggatacgggagaaggagaaggaaaccggcggaaggatcgactgggcta
 aggaaggggctgttgaggggcggaactccgacgaacaggaatgtcaggcctgccaatcagtgtcctcggtcatcatgatg
 gtgctccagtactccaacaatccagcgcacatcattgccagctcctggagtgcctgatgactcttaagcacaatgtcgtcaaggacatc
 ctctgcgttgtggcatacggaaaccgctgttcccgcacctcggctgccaagctgctcttactactggccagcctttaacgccaatc
 20 tgttcgatcgcaaagtcctactctccaaactaaccaatgacctagtgccttcacctgccaacgggagcactgtccgaactccggg
 aatgcggaggcagcaaaggtgtgctacgaccacagcattagcatcgatacgcgcccattgtccaccgcccctttacctgtgca
 tcgagtgcgccaacgagattcatcgggagcagcgaagcctggagtgcggcagacattctgcatcccatgcagcaggtatcgatgg
 tgtgcgaaaacaagaactgtcgtcccaacgagaagtccgccttctccatctgcttctccacggagtgtgccagcttcaatggcaac
 catccgatccgctactgcagccagtgcacagtaataggcacaattcccggcgaggtggcgatcacgtgggtccatcgagctgtgc
 25 agcccgctggcagatggatccagagatgcagatgcacatggtggagtgcgttggaagccttctgcgagaggcgaagccacta
 aactttgagccggcaaggagtctcgtcgtccgagtcacaaaagaacggctccggcatcacagctgacaatatttctctggagg
 aacgccagagactgggacgctatggtatctggctactggtgggtcgtgtacacccactgcagatactcccgtagaagtctggg
 caggattctgagcatgctcttccactggtttcatgtaaccgcttactcatacgtggtttatatacctgcctggtgccacatccccgga
 gtatgcccggtgttgaggccactgggagaccttggtgcgcgaacaagccacttgaaagagggtcttcagcggcttatatgcctg
 30 gtgccatatgaggttatcacttccgaaatttgggactatgtgatgccgcactggatggaggccatcaccaacgacgtggccgaga
 aggaactgaacgagctgaagattgtgctcagcaagatcctcgatccggaatgtcgcctctgggctttgatgccaaaacctgtac
 aactttgtggccattcgatttgagaagacaacggcaaaaggtgcagcagcaggcactccactggctgcagatcctcaccaagctgg
 agattctcattccactgggtccagttgttcgcatgttcggcgatggtgttcgcataatgaaatacggcatccagcacgagctgatgcg
 cgagaaggatgcccaatctcagtcctggccaaggctcccaagaccccggtgtaaagagagcaaggagaccaaaagcggatgtg
 35 gccaatccgcccaggcctcctgtgtcgaggatgactctggttaatacgtctgccatttcggatgacgaggcgcccacgaatcgtca
 cacggaattctccacggatgctgagcacaatctcacctgttcgcatcctcatgctggacatacttctgaagcaaatggaactacagga
 cgtggagcagcacatgggcatccatacagtggtctgcgagaacgtctccaggctgatcaagtgcagtggtcactgcagctcgagt
 ggtgtcagtagtcatgtctgcgccttaaagggtcccatcgaggacatcattgaggaagaaaagtctcgcgcaaatctccaccg
 aatccgacaaggaaaagacccgtgatcgagatgttccctctcgatggctccactaccattccgctgggacctttaggaggattg
 40 cagacccttaa

Human Homologue BLASTX with EST: Phosphatidylinositol transfer protein
 (P48739)

45 **Drosophila EST** SD01527 (AI530804), GH18602 (AI387906)

Annotated Drosophila genome genomic segment

AE003725

Annotated *Drosophila* genome Complete gene candidate CG5269 – vib PIP transfer protein

5 **Human homologue of Complete gene candidate** 1e-90 1346772 P48739
 PPI2_HUMAN
 PHOSPHATIDYLINOSITOL
 TRANSFER PROTEIN BETA
 ISOFORM

10 **Putative function** phospholipid transporter involved in lipid metabolism

Confirmation by RNAi Slight reduction of G1 and increase in G2/M peaks
 indicating arrest in G2/M

15

| | | |
|----|-----------------------------------|---|
| | Line ID | 418/32 |
| | Category | Meiotic defects in testis: cytokinesis defects. Dark bands in eyes, dominant. |
| | Reversion | ? |
| 5 | Map Position | 69C |
| | Rescue ID | G2E |
| | Rescue Sequence | |
| 10 | | AGCTAAATAACAAACTCATTACTAGTATATTACTGCCGCCGATTTGCAAGCGC GTACCGATCCCGATACCAGGCCAATCGCACTCCCCAGTTGTACGTCATCACTT AAGTAATAAATCAGCGGCCAAATCGCATAAATTGCTATTGATATTCCGCCCGCT GTGTGTGCGTGTGTATTTGCAAAAAGAGTGTGTGTGTGTATGTGCATATGACTC GTGCGTTTAGCCGACAATTGGAGAAAAAGCATTAGAATCCCAATTGGCTAACT AAACTAAAGTTGGCTTGGCCAAACATAAACAAAAAGTGCGGGCGCAGCGATT 15 TGGCAGCGAAACAAAAACACCAAAGTGTTATTGGCAGATATATATGTTAATTA AATATNAAAAAGTGCGTGCGAA |
| | Genomic hit, Accession No. | AC006589 |
| 20 | <i>Drosophila</i> EST | SD01527 (AI530804), GH18602 (AI387906) |

Rest of results same as line 1104/16

Example 4 (Category 1)

Line ID 1285/1
Category Meiotic defects in testis: cytokinesis defects
Reversion ?
Map Position 85D1-5

Rescue ID D8E

Rescue Sequence

10 GTTCGCAAAAAATATATCTCACCGTGAGTGCGAAAGAGAAAAAGAGAAGCGG
 AGAGGTGGAGAGCAAGTGGACATGAATCGTCGAGAGTCAGAGAGAGAGAGG
 TGGAGAGGGGTGAGCAGCTGTTGTCTGACAATAACATAATCAGCAACAATTTAT
 GCTGTTTAAAAAGAGCAAGAGAAACGCTAATGAAGGGGAACACGGGCAGGGT
 CAGGGGTGGTGGATCCCCTACATATCTCTCTCTTTACCGCCCCCGCTCTGGC
 15 ACCCTCTCTGTCGCTCTCCCATTAGCCGCACACGTGCAAGCTTAGCATTCTATC
 TGTCTGTCTCTGTTTGTGTTTGTGTTGCTAAGCCGAATTCT

Genomic hit, Accession No. CSC:AC014256

Associated ORF

Genscan ORF1 predicted sequences

>/tmp/aaaaakfaa|GENSCAN_predicted_peptide_1|702_aa

MIQRCVLLWIVCFCDLFLGLLFLKRKRNAHTPPPPQFTTYRHLLCYCFRNGEIM
 ANICLSRLSVLEEIVLLLRVPCAFYFVDYYYVPCLLSVLSESFLYHDQLKVFNRTK
 25 QQHQQQQQQQQQQQLYQQHQQQQQQHYGPPPPYFQQLHQHQQQQQQQQQQQQ
 HQQHMKFLGGNDDRNGRGGVGVGTDAIVGSRGGVSQDAADAAGAAAAAAVGV
 YVFQQRPSGGVGVGVGGVGGVPGVGAVGSTLHEAAAAEYAAHFAQKQQQT
 RWACGDDGHGIDNPDKWKNPPMNPANAAPGGPPGNGSNGGPGAIGTIGMSG
 LGGGGGGGAGGGNNGSGTNGGLHHQSMAAAAANMAAMQQAALAKHNHMI
 30 SAAAAVAAQQQHQPQQHPQQQQQQQQAQNQGHPHHLMGGGNGLGNGNG
 LGIQHPGQQQQQQQQQQQQQHPGQYNANLLNHAAALGHMSSYAQSGGSMYDH
 HGGAMHPGMNGGMPKQQPLGPPGAGGPQDYVYMGGQTTVPMGAAMMPPQNQ
 YMNSSAVAAANRNAAITTSTAKKLWEKSDGKGVSSSTPGGPLHPLQIPGIGDPSS
 VWKDHTWSTQGENILVPPPSRAYAHGGASDTSNSGNAGILSPRDSTCAKVVEYVF
 35 SGSPTNKDSSLSGLEPHLRNLKFDDNDKSRDDKEKANSPFDTNGLKKDDQVTNSN
 GVVNGIDDDKGFK

>/tmp/aaaaakfaa|GENSCAN_predicted_CDS_1|2109_bp

atgattcagcgtcggtgttcttctatggatagctgcttctgcgactgttcttggggctcctgttcctcaaacgtaaacgcaacgca
 40 cacactccccccccccccgcaattcaccactatcggcatctactttgttatgttttcgtaatggggaaatcatggctaatttgc
 cttagtcgtctttcagttttagaagaaattgtttgcttttacgcgtgccttgtgcgtttattttgttgattattattatgtgccctgtctgtgt
 ctgtgttatcggaatcttttctttaccatgaccagctcaaagttttaatcgcacaaaacagcaacaccaacagcagcagcagcagca
 gcagcagcaactctatcagcaacatcaacagcagcagcagcagcaacattacgggtccaccaccgcccactttcaacagctacacca
 gcaacaccaacagcagcagcaacaacagcagcagcagcagcaacaccagcaacacatgaagttttgggtggtaacgatgatcgca
 45 atggccgcggaggcgtcggttgacggatgccattgtaggatctcgaggtggcgtctctcaggatgccgccgatgcagctg
 gtgccgccgcagccgccgccgtcggttatgtcttcagcagcgtccatcgctggtgggtggcgtcggtggcgaggagtg

ggtggcgggtgtgccaggggtcggagccgtaggctcgacctgcacgaggccgccgccgagtagccgcccaactttgcc
 agaagcaacagcagacccgatgggcgtgcggcgacgacggccatgggatcgataacccggacaaatggaagtacaatccgc
 cgatgaatccggccaatgccgctcctggcgggtccaccgggaaatggcagtaatggtgggcccggcgccattggaaccattggc
 atgggcagcggattgggtggtggtggcgggcgaggctggcgggcgaaataatggcggtctgtgtacgaatggcggtctgc
 5 atcatcaatcgatggcgctgcagctgcgaatatggcagccatgcaacaggcgggcggttgccaagcacaatcacatgat
 cacaggcagcagccgagttgcagctcagcaacaacatcagcatccacaccagcagcatccccagcagcagcagcaacagca
 gcaggcgagcaaccaggggcatccacatcaccttatggcgcggtggcaatggactgggcaacggcaatggattgggcatacaa
 catcccgccagcaacagcagcagcagcagcaacaacagcagcagcaacatcccgccagtagaacgcgaatctgcttaacc
 atgcggctgccttgggtcacatgtcatcttatgcccacatcggtggcagcatgtacgaccatcatggtggagccatgcacccggg
 10 aatgaacggcgccatgcccagcaacagccattgggtccaccggagccggaggacccaggaactatgtctacatgggtggc
 cagaccactgtgcccattgggagccgcaatgatgccgccacagaatcaatatgaacagctctgtgttcagctgccaatcgga
 atgcagcgattaccacatccactgccaagaaattgtgggagaaatccgatggcaagggcggtatcctcgagcactcccggtggac
 cgttgcaccccctgcagatccccggcatcggggatccctcctccgtgtggaaggatcacacctgggtccacacagggcgagaatat
 attggtgccgccccctcgcgagcctacgcccattggaggcgccctccgatacttcaaacagcggcaatgcgggcatactgagtc
 15 ccgcgattcgacttgcgcaaaagtgggtgaatatgtttcagtggtcgcaccaccaaaaagatagctcgtttccggattggaacc
 gcatttgcggaatctaaagtgtgacgacaacgataagtcacgcgacgataaggagaaagcaaacctcctggttgacacaaacggtt
 tgaagaaagacgatcaggtcacaactcaaatggtgtgtcaacggcattgacgatgacaagggcttcaagtga

20 **Drosophila Gene Hit** TBLASTN of ORF1: pumilio protein (L07943)
Human Homologue TBLASTX with pumilio: Soares fetal heart NbHH19W Homo
 sapiens cDNA clone (W77820)

25 **Annotated Drosophila genome genomic segment** AE003681
Annotated Drosophila genome Complete gene candidate CG9755 – pumilio RNA

30 **Human homologue of Complete gene candidate** 1e-154 1944416
 dbj|BAA19665| (D87078)
 similar to D.melanogaster
 pumilio protein (S22026)

35 **Putative function** Putative RNA binding protein which is localised to the cytoplasm.
 Wild-type allele of pum involved in development of the abdomen
 (embryos) and of the imaginal discs (larvae or pupae), perhaps
 having a function in signal transport.

Confirmation by RNAi Only wild type profiles observed

Example 5 (Category 1)

Line ID 1389/1
Category Meiotic defects in testis:segregation defect, cytokinesis defect
(Ck-09/32)
Reversion NR
Map Position 93B4-8

Rescue ID 2C9P

Rescue Sequence 1

GTTCGGGGTGTGTGCGTGCTTGCGAGTGTGCCTGTGTGTGTGTAGGAAAGGAG
CAAGAAGCAGCAGCAGCGGCAGCAGTAGAAATAGCAAAAGGAGGCAGCAAC
AACAATAAGCTAGAGAAACCGCCAGCAGCAGCCCCCTAATAAAGAGCAGAGA
AAAAAATGAGTTCAAGTTGTGAAAGGTGTGTGCCGTTACACTACAAACTACAA
CACCACCATCAGCGGCAGCAAAGAAATACAACAACAAATACGGCAATCTCCA
GACAACGCGAATGTCGAAATTGTGTATACAATTTATTAAGAAAGCAAGAGCA
GCAACAACAATGACCAGCTGCAGTTCATCAGCGGTGTCCTCCTGAATGCCGCT
GTCGTCGTTGGTGTCTGCCACCGGCGGTTCTCAATAATAAGGGCAGGAGGAG
CTGCTTAGGTGCACACAATGTAGTTTGGCTTGGTGAATGCTTCTCTTTTGTG
CTGCTGGCGCATACGTTCTCTTCTCCCCTCATGATCTCAGTTGTCTGCATCGA
TGAGCCGCCACCAACGGTGGCTTCTTCTGCTCCTCTTTGGCAACGGACTGCTG
CAGTCTTGCCAGAATTTTTCTTAAATACTGAGCTTCAACTTGGTCTGCTTGGT
AATGGTATACCATAAGCCATGGACTTGATGCCCCTACAAAGCTCTGTGATTG
AAATGGGATGCA

Rescue ID 2C9E

Rescue Sequence 2

CCCCGAACGCACTTTATATATATAAATATATATATTATTTTCTTTCCTTATTTT
CGTTTCGGCCGCGACAGCGAATATGCAATTTTCCTCTCAATTGATTTTTTTACA
CACTCGCACTCCTTTTTCACATGCGTGCAGTTTATGTTGCTATTGCTGCTACTGC
TGCTGTTGTTGTTATTGTTGTTCTGGCTGCCGCTGCAGTGCAACTTGTAACACT
TTCACATTTATGACATAATGCACTGGCCATATTTTTGCTTGGCTCTCCGTTTGT
GCAACTGCATGTTCCCAGTGCTTTTTTAATATTTATGCTGCAGTGCGTGCAAAT
TCGAACGCGAGACGATCCGCTTTTCGCTGCATCTATGCGCTGAAGATGTGCTG
CAGTCGATGGGCTCGTCGATAGTGGGAAGGCTCGGTGCCGGCACTATCGATTC
CCAACACCATACGATAATATCGGCTAAAGTTATCAATATCGAAGTTTACTATA
TTTCGGGTTTTTACGTTTTAAATCTACCTTATCAACATTTTTTGNAAGAAGTAAA
AAGTAGTTCTCTTATGGATGCATC

Drosophila EST several including LD10379 (AA816796)

Annotated Drosophila genome genomic segment AE003733
Annotated Drosophila genome Complete gene candidate CG3421 - novel protein with
weak homology to myosin

| | | | |
|----|---|---|---|
| 5 | Human homologue of Complete gene candidate | | Ensembl predicted Gene:ENSG00000071333 Clone:AC022505 Contig:AC022505.00011 5.60E-37 (predicted protein with Core domain in kinesin and myosin motors ENSG00000087179) |
| | | | |
| 10 | Putative function | Possible novel motor protein involved in cytoskeleton organization | |
| | Confirmation by RNAi | Marked reduction of G1 and G2/M peaks indicating fewer cycling cells | |

Example 6 (Category 1)

Line ID 293/9
Category Mitotic defects in brain: cytokinesis defect
5 (no overcondensation of diploids, very high polyploidy)
Reversion NR
Map Position 66B

Rescue ID 2G5E

10 Rescue Sequence
GTACAAACGAATTATTTGTCTCCTTGTGCGTTCGTTTTATTGTGTTTCGAGTTCT
GTTGGTGTGTGTTTTTGTGTATGTTCCACGAGTTGTTTCGCATTAAAAAATTAAC
TGCAGAAGATCCATGGAAATGGAGACCATTGAAGAGCAATCGAAGTGCGGTG
AGTACTGAAAGAGGGCGCGGGGGCGTGGCAGCTCCAAATGGCCGGCGAATTTA
15 TCATTTTTCAATGTCGTCCAAAGGGGTTGGGTACGGGGTAAAACACATTTCGG
GGCCAAAAGATCCTCATAAAAAATGTCGCTGCCAGCAAATGCAAAAAATAAA
ATAAAATAAGAACGACTATAAGTACATCTTTGTGTGTATTTGTGTGACTAAAA
AAGCAACGGCATCGTGTGCGCANATATTTTAATCTTTNTTTCTGAATTTATTTTCG
GTGTACAAAATATTTATCGCATAAATGCGAAATGCCTCCCTCTCTTCATCATCG
20 T

Genomic hit, Accession No. AC008303

Associated ORF

25 Genscan ORF1 predicted sequences >20:53:38|GENSCAN_predicted_peptide_3|261_aa
MMDNDDALLNNGGPQSGAETVYGTEDNNMVMSEKCRIFPATQRTGFVGFATFSG
VLLLDLQALQHCDEVIRIDVNIATLEQIKRERQEELAAERERIRAQIAADRAEQQAQRF
NTPDISSTTNSVAATAASNVTDDASVSSVDETRLQIRLPGGIQRKTSFPAGEVLAT
VRVYVRNEMLAASDVRDFTLATSYPRREFQTEDEVKTLNELNLVPNAVVLVLT
30 EQVNPADDQTAKRSASTKRTKTHRHKRQLMADEPQSDHYKN

>20:53:38|GENSCAN_predicted_CDS_3|786_bp
atgatggacaacgatgatgcactgctcaacaatggaggaccacagtcgggagctgaaactgtctacggtaccgaggacaacaac
atggatcatgtcggagaagtgcgcatttcccgccgactcagcgtactggatttggggcgcgacgtttcgggagtgctgcttctt
35 gatcttggtgccctccagcattgtgatgtgatccggattgatgttaacattgcaacgctggaacagattaagcgtgagcgtcaggag
gagctggcggccaggagcgcattcgtgcccgaattgcagccgatcgggcagagcaggcacaacgtttaatacgcgggacat
tagcagcacgaccaattcgggtggcggccaccgctgcctccaacgtgatcacaacagacgcctcgggtgagttcgggtggacgaga
cgaggctgcagatccgactaccggtggcattcagcgcaccaaatcctttccagccggcgaggtgctggctaccgttcgtgtcta
cgtgcgaaacgagatgctggcggcgagcgtatgcgcgactttaccctggctaccagttaccacgaaggaggattccaaacgg
40 aggacgaggtcaagaccctgaacgagctaaatctagtgcctaatgcgggtggttctggtgctgaccaaggagcaagtgaatccag
ctgatgaccagacagcaaacgatcagcaagcaccacacacagacacaagcggcaattgatggcagacga
gccacaatctgaccattataaaaactga

45 Drosophila Gene Hit rescue sequence: pebble (rho1 GTPase exchange factor)
(AF136492)

Human Homologue BLASTX with pebble: KIAA0337 (AB002335)

***Drosophila* EST** SD09146 (AI542703), SD01796 (AI530981)

Annotated *Drosophila* genome genomic segment AE003557

5 **Annotated *Drosophila* genome Complete gene candidate** CG8114 - pbl pebble rho1
GTPase exchange factor

10 **Human homologue of Complete gene candidate** 2224615 dbj|BAA20795|
(AB002335) KIAA0337
[Homo sapiens (3e-21) also
mouse homologue 3e-95
42359 transforming protein
(ect2) - mouse >gi|293332
(L11316) ect2 [Mus
musculus]

15 **Putative function** A guanyl-nucleotide exchange factor involved in signal
transduction which is localised to the mitotic anaphase. pbl is
required for the formation of the contractile ring and the initiation
of cytokinesis in *Drosophila*

20 **Confirmation by RNAi** Slightly reduced G1 and G2/M peaks indicating fewer
cycling cells

| | | |
|----|-----------------------------------|--|
| | Line ID | 542/3 |
| | Category | Mitotic defects in brain: cytokinesis defect (very high polyploidy) |
| | Reversion | NR |
| 5 | Map Position | 66A |
| | Rescue ID | 2A1E |
| | Rescue Sequence | |
| | | GTCCAGTTAATGAAAGTAAACGAATCGAGTACAAACGAATTATTTGTCTCCTT |
| | | GTGCGTTCGTTTTATTGTGTTTCGAGTTCTGTTGGTGTGTGTTTTTGTGTATGTT |
| 10 | | CCACGAGTTGTTTCGCATTAAAAAATTAAC TGCAGAAGATCCATGGAAATGGA |
| | | GACCATTGAAGAGCAATCGAAGTGCGGTGAGTACTGAAAGAGGGCGCGGGGC |
| | | GTGGCAGCTCCAAATGGCCGGCGAATTTATCATTTTTTCAATGTCGCCCCAAAGG |
| | | GGTTGGGTACGGGGTAAAACCATTCGGGGGCCAAAAGATCCTCATAAAAAA |
| | | TGTCGCTGCCAGCAAATGCAAAAAATAAAATGAAATAAGAACGACTATAAGT |
| 15 | | ACATCTTAGTGTGTATTTGTGTGACTAAAAAAGCAACGGCATCGTGTCGCANA |
| | | TATTTTAATCTTTTTTTTCTGAATTTATTTTCGGNGTANAAAATATTTATCGCATA |
| | | AATGCGAAATGCCTCCCTCTCTTCATCATCGNTTCCCCTNACTCTCCCTCTCTT |
| | | CGCCCGACACTGTACCGACGCAAGAAGAAC |
| 20 | Genomic hit, Accession No. | CSC:AC018042 |
| | Drosophila EST | SD09146 (AI542703), SD01796 (AI530981) |

rest of results same as line 293/9

Example 7 (Category 1)

| | | |
|----|---|---|
| 5 | Line ID | 229/30 |
| | Category | Mitotic defects in brain: cytokinesis defect. Meiotic defects in testis: cytokinesis defects (Mitotic higher level of condensation, polyploidy, Meiotic: |
| | | Ck05/07) |
| 10 | Reversion | ? |
| | Map Position | 91F |
| | Rescue ID | A7E |
| 15 | Rescue Sequence | |
| | | TCTTGGCCAAACAACGCGAGCAGCTGATGTCGCATGGTGGGAAAATGAGGGT |
| | | GGCGCGAGTGGAAGTTGCCATATCGCTGCGATCACAAGCAGCAAATATGGAA |
| 20 | | GATTAAGCGGAAAACGAAAGACAAAATAATTACAATCAAACAACCGAATTAT |
| | | AAAAAGAAAATGGTTTGTCTCCGAGTTCGTTTAAATATGCTTATCTACGTATC |
| | | AATTAAAAAAACCGTAGAAAGAAATTCACGATTCACCCTAATCTAGCTAAGA |
| 25 | | CACCAACCAAAAATTTCCGATTTACTTTCAGTTGAAGTTGTTGTTACACACTTT |
| | | TCTTGTCGATGTTTTGAAGCGCCCATTTGAAATTGATCATTTGAATGTTTTTCCA |
| | | AATTACCCACATCCATTACAATAAATTTAAATTGCTTATTATTTGATTTTACT |
| 30 | | TGGGAAAATCCCGTTGCCAAATTGGAATTACAATTCCAGCTTGGAATCCGTCA |
| | | AACTTTACAACATAAACTTATTGTTCTTTTCCGGACAATGCTTCCA |
| | | |
| 35 | Annotated <i>Drosophila</i> genome genomic segment | AE003686 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG6284 - novel protein possible sir2 family of transcriptional regulators/telomeric silencing |
| | Human homologue of Complete gene candidate | gi7706710 0268A424791DE5BF [ref]NP_057623.1 sir2-related protein type 6 [Homo sapiens] (1.10E-74) |
| 40 | Putative function | Putative transcriptional regulator |
| | Confirmation by RNAi | Complete loss of G1 and G2/M peaks indicating fewer cycling cells |
| | | |

Line ID 1104/16
Category Mitotic defects in brain, Cytokinesis defect (no overcondensation of diploids, high polyploidy)
Reversion R
5 **Map Position** 92A

Rescue ID B5E
Rescue Sequence
10 GTCCGGAGCGGAGCTAAAGTTCGATGTTTCGTGCAAAACACTTCGATTCCGATA
GGCGGATGCTATCGATTTTCGGCGATGCCCCGTTGGTCACACTTGGTGGTGGGCG
CTGCCCCGTCGCCGACTATCGATAGCACAAAGCGGGTTATTTAGGTGTGCGCAGC
TTGTAAGGGTGACTCATGCTGTAAAATTATTATAAAAAGTTAATGAATATAA
TATAGTTATAATAAAATTATATATAAATCTATAAGATCAAAGATCATCAGTTA
TCATTTATCATTTGATTATATGAAAAACAAGAACAGAAACAAGATTTAATAGG
15 TTTTGTGAAATGTGAAAATGTGGGTTACCCCCAATTCTTATTCGAAATTAAATAA
CCTAAAGAACAGTTATACACAGATAGGTAATTTGCACATAAGCCAAATTTTGT
CTAGAATTCCGCGGAATTAATTCTTGAAGACGAAAGGGCCTCCGTGATACGCC
TATTTTATAGGTTAATGTCATGATAATAATGGTTTCTTA

20 **Rescue ID** B5P
Rescue Sequence
CTCCGGACACGCAGTAGCTAAATAACAAACTCATTACTAGTATATTACTGCCG
CCGATTTGCAAGCGCGTACCGATCCCGATACCAGGCCAATCGCACTCCCCAGT
TGTACGTCATCACTTAAGTAATAAATCAGCGGCAAATCGCATAAATTGCTATT
25 GATATTCCGCCCCGCTGTGTGTGCGTGTGTATTTGCAAGAGAGTGTGTGTGTGT
GTGTGCATATGACTCGTGCGTTTAGCCGACAATTGGAGAAAAAGCATTACCAA
TCCCAATTGGCTAACTAAACTAAAGTTGGCTTGGCCAAACATAAACAAAAAGT
GCGGGCGCAGCGATTTGGCAGCGAAACATATACACCAAAGCGCTATTGGCAG
ATATATATGTAGATTAAATATAGAAAGTGCGTGCGAAGGTTAAGAGTCGAGT
30 GCAAGTGCATTTATATTTGGAAATAATAAATGCTACAAT

other results same as 229/30

Example 8 (Category 1)

Line ID 343/5
Category Mitotic defects in brain: cytokinesis defect
5 (very high polyploidy, chromosomes entangled?)
Reversion NR
Map Position 75B

Rescue ID C6E
10 **Rescue Sequence**
GGTTTCGAGTTCGTTTCGGTTTCGGCCTCTCCGTTTCGGCTCTCTCTCGCCATCCC
GCTGCCGCACACATTGGCCTCTCTCTCGCAGCTCCACATTCGAAGGTGGCTGA
CCGAAATGTGGGTCACGACAATGGCGGGGTTTCGTTGAACTGAACCACCGCCG
CAGTCGCTGCCGTGCTCGCTGCTCTGCCTCTGCTGACGTCGTTAACGTTTTGGG
15 GCTTTCGGTTACGTAGCTCGTGTGCGAGCGAGAGGGGCTACTAGAGGGACTGC
GACACACAAGTTGTGTGCATTTTTTGGCCCCAAAAAATCACAATGGGACACAAA
ATATTATTTAATACATCACATAATTGTTTAATCATCTGGCTGGAAAGTGTTCGAG
TTCATCGAACTGCCAGCGATTGACAAATTGCGATTTTCAATGCGGCACAAAATA
TTTACTCAAGCAAATTGTTTGCCTTCGTTAATTAGGCGGGGAGTGCCGCCAA
20 ATGCGGTCATATTGCAGAAGTATCCAAGAAGTTGGAGAAACAAGCTGCTTAA
ACATTAATTAACACACACCTAAATGGATACATTTGCTACAAACAATTATAAAT
GTTACCCTTATATTAATTTTCAAATTTCTAAATAATCAA

Genomic hit, Accession No. CSC:AC015427
25

Associated ORF
Genscan ORF1 predicted sequences
MVCAMQEVA AVQHQQQQQQQLQLPQQQQQQQQQT TQQQHATTIVLLTGNGGGNL
HIVATPQQHQPMHQLHHQHQQHQQHQQQA KSQLKQQHSALVKLLESAPIKQQ
30 QQTPKQIVYLQQQQQQPQRKRLKNEAAIVQQQQQTPATLVKTTTTSNSNSNNTQT
TNSISQQQQQHQIVLQHQQPAAAATPKPCADLSAKNDSESGIDEDSPNSDEDCPN
ANPAGTSLEDSSYEQYQCPWKKIRYARELKQRELEQQQT TGGSSNAQQQVEAKPA
AIPTSNIKQLHCDSPFSAQTHKEIANLLRQQSQQQQVVATQQQQQQQQQHQQHQQ
QRRDSSDSNCSLMSNSSNSSAGNCCTCNAGDDQ QLEEMDEAHD SGCDDELCEQH
35 HQRDSSQLNYLCQKFDEKLD TALSNSSANTGRNTPAVTANEDADGFFRRSIQQK
IQYRPCTKNQQCSILRINRNRCQYCRLKKCIAVGMSRDVLRLEQPKAGAKNKSCE
PSKNSTVNQINSKLELGNSNEMK

>21:55:09|GENSCAN_predicted_CDS_1|1533_bp
40 atggtttgtcaatgcaagaggttgctgccgtgcagcatcagcagcagcaacagcaactccagttgccccagcagcaacagcag
cagcagcagacaacacagcagcaacatgcaacaactatagtgtgctgacgggcaatggcggcggtaatctgcacattgtcgcc
acaccgcaacagcatcagccgatgcacagctccaccatcagcatcagcatcagcatcagcaccagcagcaggccaagagcc
aacagctgaagcaacaacactcggcgctggtcaagttgctggagtcggcgcccatcaagcagcaacagcagacgcccgaagca
aatgtttacctgcagcagcagcagcagcaaccgcaacgcaaaagactgaaaaacgaagcagcaatcgtacaacagcaacaac
45 aaacacctgcaacactagtaaagacaacaaccaccagcaacagcaacagcaacaacaccagacaacaatagtattagtcag
cagcaacagcagcatcagattgtgtgcagcaccagcagccagccggcgagcaacaccaaagccatgtgccgatctgagcg

ccaaaaatgacagcgagtcgggcatcgacgaggactccccaacagegatgaggattgcccgaatgccaacccggcggggcac
atcgctcgaggacagcagctacgagcagtatcagtgcacctggaagaagatacgctatgcgcgtgagctcaagcagcgcgagt
tggagcagcagcagaccaccggaggcagcaacgcgcagcagcaagtcgaggcgaagccagctgcaatacccaccagcaac
atcaagcagctgcactgtgatagtcccttttcggcgcagaccacaaggaaatcgccaatctcctgcgccaacagtcccagcaac
5 aacaggttgtggccacgcagcagcagcagcaacagcagcagcagcaccagcaccagcaacaacgaagggatagctccgaca
gcaactgctcgctgatgagcaactcgagcaactccagtgcgggcaattgttgacctgcaacgctggcgacgaccagcagctgg
aggagatggacgaggcccacgattcgggctgcgacgatgaactttgcgagcagcatcaccagcgcactggactcctcccaactg
aattacctgtgccagaagttcgatgagaaactggacacggcgctgagcaacagcagcgccaacacggggagggaacacgccag
ctgtaacagctaacgaagatgccgatggattcttcgccgctccatccagcaaaagatccagtatcgcccgtgcaccaagaatca
10 gcagtgcagcattctgcgcatcaatcgcaatcgttgtcaatattgccgcctgaaaaagtgcattgccgtgggcatgagtcgcgatgt
tctgcgcctagagcaacctaagctggtgccaaaaataagtcattgaaccgagcaaaaattcgaccgtcaaccaataaacagc
aaactcgaactcggcaacagcaatgaaatgaaatga

Drosophila Gene Hit TBLASTN with ORF1: ecdysone-inducible gene E75B (X51549)
15 and nuclear receptor superfamily protein (U01087) BLASTN with
genomic sequence matches ecdysone inducible gene

Annotated Drosophila genome genomic segment AE003522
Annotated Drosophila genome Complete gene candidate CG8127 Eip75B ecdysone-
20 inducible gene E75B nuclear
receptor NR1D3

Human homologue of Complete gene candidate ORPHAN NUCLEAR
25 RECEPTOR NR1D1 (V-
ERBA RELATED PROTEIN
EAR-1) (REV-ERBA-
ALPHA) Q15304 (9.40E-74)

Putative function Ligand-dependent nuclear receptor, putative transcription factor
30

Confirmation by RNAi Slightly reduced G1 and G2/M indicating fewer cycling
cells
35

| | | |
|----|-----------------------------------|---|
| | Line ID | 448/23 |
| | Category | Mitotic defects in brain: cytokinesis defect (very high polyploidy |
| | Reversion | NR |
| 5 | Map Position | 75B |
| | Rescue ID | 2G4E |
| | Rescue Sequence | |
| 10 | | GCTGGTGGACGCTGCTTTCATTCGCAAATTGCTCGTCGTTGGCAGCGGTTGTGC AGAGCAAGAAAAGCGCGCGGAAAAACCAAGCAAAAAATTAATACAGCTGGAT CAAGCGAAAGAGATAGAGAGCAGAGTCAACAGCAACAAATGTTCAATAGCA AATGATATCGCATATTTTTTGTGTTGGTGCCAGTGAAGTGAGATCAAAGTGAAGTG TGCAATGTTTCCTTATTAGCAAATCGTAGAGCAACCAACAATCGAGAGTTCAAG TGTCATTTTCGAAGCCAAAAAGCAAAATCTCTAATTCAAATATGGTTTGTGCAA 15 TGCAAGAGGTTGCTGCTGTGCAGCATCAGCAGCAGCAACAGCAACTCCAGTT GCCCCAGCAGCAACAGCAGCAGCAGCAGACAACACAGCAGCAACATGCAAC AACGATAGTGCTGCTGACGGGCAATGGCGGCGGTAATCTGCACATTGTCGCCA CACCGCAACAGCATCAGCCGATGCATCAGCTCCACCATCAGCATCAGCATCAG CATCAGCACCAGCAGCAGGCCAAGAGCCAACAGCTGAAGCAACAACACTCGG 20 CGCTGGTCAAGTTGCTGGAGTCGGCGCCCATCAAGCAGCAACAGCAGACGCC CAAGCAAATTGGTTACCTGCAGCAGCAGCAGCAGCAACCGCAACGCAAAAGA CTGAAAAACGAAGCACAATCGTACAACAGCAACAACAAACACCTGCAACAC |
| | Genomic hit, Accession No. | CSC:AC015427 |
| 25 | Drosophila EST | GM03519 (A801874) |

Other results same as line 343/5

Example 9 (Category 1)

| | | |
|----|--|---|
| | Line ID | 36/1 |
| 5 | Category | Meiotic defects in testis: cytokinesis defects (Ck-04/06) ` |
| | Reversion | R |
| | Map Position | 79C |
| | Rescue ID | A8B |
| 10 | Rescue Sequence | GAGTAAAGTAAACTACAGAGAAAAAACGCTTTACGGCGAGAGAACGCTTTAA TTATACTTAATTTGTTGTTAATCAAACGCACAGAGCACACAACACAGAAACAC AAAACACCCGCTTGGGAAAAATCTGTAGGTAGANGAAAGGAGCTCACGTTTTT CTGGTGCAGATCGAAATCGGTATCGGGTTTATTCGCTTTGCCGGATTGTTACTT 15 CACGTTTGTTAATTGCGTTTCTTTGTTTCTTATTCTCCTGCGCACACTTTGATTT GCGTTTGCAACTCGCAATTCGCAATTGGCATTGCTATGCGACAACCTGCCGTT ATTTCCGGTCCGTTTACTTTTCCAATGGCTTGCCTACACACCGCCAAACTTTGG CTTGAACCTGGGATATTGGTTGCCCCGAATTTCCTGANAAATTTTTCCTT |
| 20 | Genomic hit, Accession No. | CSC:AC013886 |
| | Associated ORF | Genscan partial ORF1: >18:33:59 GENSCAN_predicted_peptide_1 99_aa CICFALLGLLIRRKLMVVFGSTSRKAQSLESRRAKNTSQQIGNQYPKFSQVCGKPS 25 KSNDNRNGSCRANANCELRVANANQSVRRRIRNKETQLTNVK |
| | | >18:33:59 GENSCAN_predicted_CDS_1 300_bp tgtatctgcttcgccctgcttgggctactcatteggcgaaaattaatgggtggtgttcggttctacgtcgcgcaaggcacagtctctaga gtctcgcagagctaagaatacatctcagaaaatcggcaaccaatatcccaagttcagccaagtttgcggcaagccatcgaaaagt 30 aacgaccgaaataacggcagttgtcgcatagcaaatgccaatgcgaattgcgagttgcaaacgcaaatacaagtgtgcgcagg agaataagaaacaaagaaacgcaattaacaaacgtgaagtaa |
| | Drosophila Gene Hit | rescue sequence and TBLASTN with ORF1: nucleic acid binding protein (mub) (X99340) |
| 35 | Human Homologue | BLASTX with nucleic acid binding protein: poly(rC)-binding protein 2 (hnRNP-E1) (S42471) |
| | Drosophila EST | several including LD32520 (AA951799 BLASTN matches nucleic acid binding protein (X99340) |
| 40 | Annotated Drosophila genome genomic segment | AE003596 |
| | Annotated Drosophila genome Complete gene candidate | CG7437 - mub mushroom bodies RNA binding protein |
| 45 | Human homologue of Complete gene candidate | 4826886 ref NP_005007.1 pPCBP2 poly(rC)-binding protein 2 |

>gi|542853|pir||S42471 (4e-75)

5 **Putative function** A putative RNA-binding protein specifically expressed in the CNS
 of *Drosophila melanogaster*

10 **Confirmation by RNAi** Only wild type profiles observed

Line ID 472/22
Category Female sterile
(anaphase bridges, lagging chromosomes)
Reversion ?
5 Map Position nd

Rescue ID sau 5'spl

Rescue Sequence

10 GCACGATCNCTAAAGTCTNGCANAGCTAAAAATACATCTNAGAAAATCGGCA
ACCAATATCCCAAGTTCAGCCAAGTTTGCGGTGTGTAGGCAAGCCATCGAAAA
GTAACGACCGAAATAACGGCAGTTGTCGCATAGCAAATGCCAATTGCGAATT
GCGAGTTGCAAACGCAAATCAAAGTGTGCGCAGGAGAATAAGAAACAAAGA
AACGCAATTAACAAACGTGAAGTAACAATCCGGCAAAGCGAATAAACCCGAT
15 ACCGATTTTCGATCGGTGCGGGCCTCTTCGNTATTACGCCAGNTGGCGAAAGGG
GGATGTGCTGCAAGGCGATTAAAGTTGGGTAACGCCAGGGTTTTCCAGTCACG
ACGTTGTAACGACGGCC
ANTGCCAAGCTCTGCTGCTCTAAACGACGCATTTTCGTACTCCAAAGTACGAAT
TTTTTCCCTCAAGCTCTTATTTTCATTAAACAATGAACAGGACCTAACGCCNGT
20 AAC

Rescue ID Sau 5'splac

Rescue sequence

25 GTTGTGATCNTCTTGGTNAATCENNNTTGGAAATTCCCCTAANGCTTCCGACAA
TGACCCNGNCNTACNNAGCAAANAATNGNAGNACNNGCNGNTGGNCGTANT
ANCAANAACAGGCCCGCACCGATCGAAATNGGNATCGGNTTTATTTCGCTTTGC
CGGATTGTTACTTCACGTTNGTTAATTGCGTTTCTTTGTTTCTTATTCTCCTGCG
CACACTTTGATTGCGTTTGCAACTCGCAATTCGCAATTGGCATTGCTATGCGA
30 CAACTGCCGTTATTTTCGGTCGTTACTTTTCGATGGCTTGCCTACACACCGCAAA
CTTGGCTGAACTTGGGATATTGGTTGCCGATTTTCTGAGATGTATTCTTAGCTC
TGCGAGACTCTAGAGACTGTGC

Other results same as line 36/1

35

Example 10 (Category 1)

| | | |
|----|---|--|
| 5 | Line ID | 459/2 |
| | Category | Mitotic defects in brain: cytokinesis defect. Meiotic defects in testis: cytokinesis defects: (mitotic: high polyploidy, no diploids, higher mitotic index, meiotic: Ck-01/05) |
| | Reversion | NR |
| 10 | Map Position | 66B1-6 |
| | Rescue ID | 2D5P |
| | Rescue Sequence | GCTCCGTTTCGAAAGTTGAGAGAGACTTGAAACATATGTTTCGGCGTTGCTAGAG CTGGTCGGCTACCGATAGAAACATCGATAGGTCCGATGTTTTTTACTCGTATAT 15 TGATTCANAGTTTGGCTATCGATGTTTTTAGAGTGCCCGCACATTATCTATTTT CATCTCTATTTTCGTTGGTATTTTTTTGTATTTTATGACATTTTCGACTGCAAAAGC AGGATGGCAACGCCAGATTGCCGCGAAAGTACGTTATTTTTTAAATTGGCGCAT TGAATATGAAAAATTGCAGGCACATACAGTTTCTAATAAATAATAGCAATAAT TATTATTAGCTTGTATCATACGAAGTGCACATTACAGCTACGCATCTGAAAT 20 AATAATTTTAATATATCGTCTTTTCTCCCATCGATAGAGTTCCGCGCCTATCGA TATATCGTTGATCACCAAATAAATAAAACTAAATAACGCCGCAATGGAACAC GCGACGAGTGAATTGAGGGAATTTATCTCAGATCTTGTAATTCCGCACCACGT TGCAATGGTAACATCAATCCGGATCACATCACAATGCTGGAAGGCACCCAGA TCCAGAACAG 25 |
| 30 | Annotated <i>Drosophila</i> genome genomic segment | AE003557 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG8038 - novel gene ribonuclease P homology CG7892 nmo - protein serine/threonine kinase involved in eye morphogenesis |
| | Human homologue of Complete gene candidate | CG8038- 5e-24 4309676 gb AAD00893 (AF001176) ribonuclease P protein subunit p29 [Homo sapiens/ |
| 40 | | CG7892- protein kinase mitogen-activated 7 (MAP kinase)' gi:4506093 and gi7706445 D919050533B3C33A |
| | | [ref NP_057315.1 nemo-like |
| | | |
| 45 | | |

kinase [Homo sapiens]
(3.30E-174)

5 **Putative function** CG8038: tRNA processing enzyme Ribonuclease P protein subunit
 CG7892: a protein serine/threonine kinase involved in cell cycle,
 possibly targeted to cytoskeleton

10 **Confirmation by RNAi** Both showed a marked increase in G1 peak indicating arrest in
 G1

Example 11 (Category 1)

| | | |
|----|---|--|
| | Line ID | 623/8 |
| 5 | Category | Meiotic defects in testis: cytokinesis defects |
| | Reversion | ? |
| | Map Position | 37E1-3 |
| | Rescue ID | 2E2E |
| 10 | Rescue Sequence | |
| | CTACGGGCATTTCGCATGTTCTGAACATCTGGTGTAACAAGTTCTGAGCAGTGT | |
| | TGCCAACTCTTCAGTTAAACAGTTAAAAATAGCTAAAAAATGTTGACGGTAGC | |
| | TAAATTATAAAGCTAGAAAAGAAATGATATATGATAAAATAAGTATTTTCGACT | |
| | CACAGCATTTATTATTTAAGACGGTCAGATGAAGTTACAAAAATCCTAAATTG | |
| 15 | GCCCGCTGTATCTAAGAATTAATACCAAGAAGTTGTCATCAAAGGTCGAACTC | |
| | GACGGAAATTCTACTTTGAGTTTTTAAATTTAATAAATATGTATTTAAAATTAT | |
| | GTAAATTTGTTTGTAACAAAAATAGTATATAGTATAGTAATAGTAGTTAAG | |
| | TAGTTTTAAAAATGGCCAGATCAAAGACTTTTGAGATATGATACTAATCAAAA | |
| | GTCGAATTCGCGGAATTAATTCTTGAAGACGAAAGGCCTCGTGATCGCCTATT | |
| 20 | TTTATAGGTAATGTCATGATAATAATGGTTTCTTAGACGCAGGTGGACTTTTCG | |
| | GGGAAATGTGCGCGGAACCCCTATTTGTTATTTTCTAAATACATTCAAATATGT | |
| | ATCCGCTCATGAGACAATAACCCTGATAAATGCTTCAATAATATTGAAAAAGG | |
| | AAGAGTATGAGTATTCAACATTTCCGGGCGCCTTATTCCTTTTTTGGGCGGCAT | |
| | TTGCCTTCCTGTTTTTGTACCCAGAACGCTGGTGAAAGAAAAGATCTGAAGA | |
| 25 | CAGT | |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003662 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG17559 dnt - doughnut protein tyrosine kinase |
| 30 | Human homologue of Complete gene candidate | Homo sapiens RYKreceptor tyrosine kinase GDB:21773 |
| | Putative function | growth factor transmembrane receptor protein tyrosine kinase involved in cell growth and maintenance |
| 35 | | |
| | Confirmation by RNAi | Only wild type profiles observed |

Example 12 (Category 1)

Line ID 629/14
Category Meiotic defects in testis: cytokinesis defects
(Ck-06/09)
Reversion NR
Map Position 64D

Rescue ID 2A9X

Rescue Sequence 1
GACGGGAGGAAGTAAGTGGGAGGAGAGAGTAGTGCCTCTTTTTTACTGGAGA
AATGGACAAACTCTGGGAACTGCGAACTGCGAACTAACCGAGGCAAAAATTG
AGAAGCGAGCTGAAAGCGGAATTCAAACAACGCAGCGCTGACGGCGACGCCG
GCAGAAGCAGCGCCGCACAAGGCATGCGCACAGAGAGTAAGAAAGAGCGCG
GCTAATGAATGAATGAACGAGGCGGAATGCGGGAAAGAGCGCAGAGAGGCGC
AATGACAAAATAGTTGTAGAAAAGCGCCGGCAAGCGGAACTCCACACTCTTT
CTCACTCTCTTTCCACCCACACCCCTAGTTCACCGGAAAAAGAAAATTCGTT
TGCGGCGGGGGTGTATTTTTCACCAAAAAGAGAGTGTGTGCAAAACGCTAGA
GAGAGAGAGAGAGAGAGAAAGAACTGACGTCAGTTCTGCCTCCGTCGACGCC
GCTGCCGGCGTCCCAAAGCGCCACCACCCAAAAAAACGCGAGAAGAAGCAGA
ACAAACACACACAAAAATTCGCACAGTGGAGCAGAAATCAAGC

Rescue ID 2A9E

Rescue Sequence 2
CTCCCGTCGTTTTGAGATCAGCTGCTCTCGCAACAACAACAATACTGTA
GTTACCGTCTCTTTTGCATCGTTTCGTTTTTCGTTTGTGTCGCCAAGTGATTGTGT
GTGTGCGTAAGCTTAAAGCTGACTAACAAAACGAAACAAGAAAAAATATAAA
TTATAGGAAAATTGTTAAATTATAACCAGAAAGAGAGCGGCACTTACGTGTGT
TATTGTGTGCGTGTGCTTTAAAAAGATATAAAAATAGCAATAGAAAGTTATTA
AAGCGTTGGCAAAAAAGTCCAACGAACAGCGAGAGGAAGCGGAGAACGAAA
TAGTTAAAGCCAAAGTCGCTGCCGACGTCGCACTTGAAAACGTCGCAAAAGTT
TGTAACACACACCAGTGTGTGTTTCGTGTGTGTTTTTGCCGGCGTGCCAGTGTGCG
TGCGCCTAGAAAAGAGTAAAGAAGCAGAAGAAAAGGAAGAAGCCGAAGAAG
CAGCAAAAGAAGCCGACAGCAAAAAGTAAATAAAATCAAATGCCCCCTGGCA
GAATAATATTAAATTAAGACACATACTCAAATTAATAAC

Genomic hit, Accession No. CSC:AC015076

Drosophila EST LP08767 (AI295205)

Annotated Drosophila genome genomic segment AE003567
Annotated Drosophila genome Complete gene candidate CG10668 - novel with
homology to ssDNA/RNA
binding proteins
Human homologue of Complete gene candidate CG10668 - 3e-12 4506449

ref[NP_002889.1|pRBMS2|
RNA binding motif, single
stranded interacting protein 2
>gi|1082

5

Putative function Possible single stranded DNA/RNA binding protein

10 **Confirmation by RNAi** Slightly increased G1 and reduced G2/M indicating G1
arrest

Example 13 (Category 1)

Line ID 653/12

| | |
|-----------------|---|
| Category | Meiotic defects in testis: segregation defects, cytokinesis defect (Ck-07/35) |
|-----------------|---|

| | |
|-----------|----|
| Reversion | NR |
|-----------|----|

Map Position 75B

Rescue ID I5E

Rescue Sequence

GTAAAAGCTTAGCCCATGGCGTCGACGTCGACTGCGACAGCGACGCTAGCCG
AGGCAGTGACTGCGACGTTGGCCACTTTTCGCCTTCGTTTCGCTGTCGTTTTCA
GTTGTCTCTCGTTGCTCAAAGCGCGCGGCACGCGAACGCTCTGAAATCCCAAG
TTACAACAGCAACATCAAGCAGCAGCAACAACAGTGATTCGCTGGCAAACAA
ACAAACAAACCAACATATTTTTTGTGTATCAATTGTCGGCCTAAAACTTCACAT
AAAAGTGCGTTCAATACGAAACAAATATATTTGTATATATAGAGAGCGAAGC
AATCGGTTGCATAAATTGAATTCCGTTCAATACTTCAATATAAATATTATTAA
GTACTACAATTTGAAAACATCTTTAAATATACAACATATTTTGAATTAAGTTTA
TTTTTTTTTTTAGCCACATAGAGACATCTTTGTGGCATGCTAAATTCTGTAGTA
AAACTTTCTTGGGGAAAGTGAAAGCCACGTATCAGACCAAAATCCACCCAAC
CCTGCACACACGCATCCCCATAAAGAACGACCTTGAGCT

Genomic hit, Accession No. CSC:AC014071

Associated ORF

Genscan ORF1 predicted sequences >16:36:33|GENSCAN_predicted_peptide_2|477_aa
MLILMRPSIKLAANQNAIKAPNGPKNFLDKVLVVRWLSVCLLENHIAVTASGS
NNNNNSNNINLNLKANYQMSATSIRDSFATILLDAQNRVQNATVAAKNFMLPLR
LRSDTSGDTSNNNENNSRRARQAYNCGVNWLTTHRPKRRRQVHPPLGSTPSCNN
NSSKISRNSSSSSSNNIASATATRIFLGTSAILAIDFDNTRVPGYYQPTGEWIWVSKS
MIKQLFAVAATADDVAAAAASRGNALTFLPGKEKGPRKKAEGCGMEWSGVIEWS
GGDVMCVLSSVATVDDDDHHGGGHFDGLLGTPSALIRLNCLINPKKMRMDFEVE
VAWQIARAADLRLISMHLNVPYEMKTMKTMESVIDGGSLYQPTALFGSLFCLVY
SSAADVLLLLANCKSLAHGVDVDCSDASRGSDCDVGHFSPSFRQSLSLVAQS
ARHANALKSQVTTATSSSSNNSDSLANKOTNOHIFVYOLSA

>16:36:33|GENSCAN_predicted_CDS_2|1434_bp

atgttgatcctaattgcggccgtcaatcaaattggccgcaaataaaatgcaattaaagcgccaaacggggccgaagaacttttggg
caaagtctgtgtgtccgctgttggctgtctgtctgtctgttgagaatgggcacattgctgtcactgccagcggcagcaacaaca
caacaacagcaacaacatcaacctcaatttgaaagccaactatcaaatgtcagctacaagcatccgagattcgttcgccacgattct
tctagacgccccaaaatcgagtgcaaaacgcaactgttgcgtgccaaaaacttcatgttgccgctgcgcctgcgcagtgaaccagc
ggtgacaccagcaacaacaacgaaaacaacagccggagagcaaggcaggcttataattgtggcggttaactggttgacaacgc
cgcccgaagcggcggcggaagtgcacccgccttgggttcaacgccagctgcaacaacaacagcagtaaaatcagcagaa
acagcagcagcagcagcaacaacatcgcatcagcaacagcaacacgcatttttcttggcacttccgcgattctggccatcgacttc
gacaatacacgagtaccggggtattatcagccaactggggagtggttgggtatccaagtccatgattaagcagctgtttgctgtt
gctgccactgcggatgatgttgcgtgctgcagcttcacgcggcaatgcgttgaccttttgcgggaaaggaaaagggggccaa

ggaaaaaggcggaagggtgtggaatggagtggagtggagtggagtggagtggcgatgtgatgtgtgtgctctcgagtgtg
gccacagttgacgatgatgatcatcatggtggcggccactttgacggccttggtgggaacaccttcagcgctcatccgacttaactgc
ttaatcaaccgaagaagatgaggatggactttgaggttgaggttgcattggcaaattgctcgagctgctgatctgcggctgatctca
atgcaccttaatgtgccttatgaaatgaaaacgatgaagacgatggagagcgtgatcgatgggtggctccctgtaccaaccgactgc
5 tctcttcggttctttgtttgcttggtgtattcttcagctgctgatgtgtgtgctgctggcgaactgtaaaagcttagcccatggcgctc
acgtcgactgcgacagcgacgctagccgaggcagtgactgcgacgttggccacttttcgccttcgttcgctgtcgttttcagttgtc
tctcgttgctcaaagcgcgcgccgacgcgaacgctctgaaatcccaagttacaacagcaacatcaagcagcagcaacaacagtga
tctcgttgcaaacaacaacaacacacatattttgtgtatcaattgtcggcctaa

- 10 **Drosophila Gene Hit** rescue sequence, ORF1 and genomic sequence: Canton S E78B
nuclear receptor superfamily protein (U01088)
Drosophila EST LP11082 (AI296953 similar by BLASTN to U01088)
- 15 **Annotated Drosophila genome genomic segment** AE003593
Annotated Drosophila genome Complete gene candidate CG18023 - Eip78C
Ecdysone-induced protein 78C
nuclear receptor NR1E1
- 20 **Human homologue of Complete gene candidate** CG18023- 4e-32 119100
P20393 EAR1_HUMAN V-
ERBA RELATED PROTEIN
EAR-1
>gi|1082832|pir||A32608
- 25 **Putative function** ligand-dependent nuclear receptor , putative transcription factor
Confirmation by RNAi Not done due to failure of PCR procedure

Example 14 (Category 1)

| | | |
|----|--|--|
| | Line ID | 876/2 |
| | Category | Meiotic defects in testis: cytokinesis defects |
| 5 | Reversion | ? |
| | Map Position | 73A |
| | Rescue ID | 2H1E |
| | Rescue Sequence | |
| 10 | GATCAAACAGAAAATCCAAAAACGAACAGCGCGCGGCGAACGAGAGCCGTT | |
| | GAAGCCGGCAGAGAAGTGCGCTGCTCGCGTCGCTGCCGGTATGTGCGTGTCTG | |
| | TGCACTGAGAGAAAATGCTCGATTAAACAGAGAAAATTAATAGTAATATAAAA | |
| | AAAAAAAAAAATTTGTTTATTATTCTCAATTCAATAAAAATGTAATTATTTATTAT | |
| | ATTGGTTGTATAAGAATTTTTATAAAGTAGTATAAATTTTCAATCAAATAAAT | |
| 15 | ATGTACATCTAACAAAAAATGTTATTATCTTATAACAAAGAGGTAAAATCATA | |
| | AGTAGTACGAAATCTTTAAAAGAGAAAGTGTGTTACGCAAAAAGTATTCAAA | |
| | GCAGTCTTTTATTTAATTTAATTTAATTTATTTGTGCTTTATCCCTTATATATATA | |
| | TGTACATTTTCATTAAAGCTAATGGTATAATTAGGTATTTACAGTGTTTAGCTAA | |
| | GGCTTTCATCTGAAATATTTATTAATTATGTCTAGTTGACCTGTTTTTAGTTTTT | |
| 20 | TTGNATAACAATATTTATTATTTATTAAGGAAAACAAGGGGAGAAGAAAAAC | |
| | CTTAATTGAAGCAAAGCAGTCTTTTGAACCCACTGGTG | |
| | Genomic hit, Accession No. AC005633 | |
| | Drosophila Gene Hit rescue sequence: argos (M91381 | |
| 25 | | |
| | Annotated Drosophila genome genomic segment | AE003527 |
| | Annotated Drosophila genome Complete gene candidate | CG10162 – Egf2 translation facto |
| 30 | Human homologue of Complete gene candidate | CG10162 - 4e-11 181969 (M19997) elongation factor 2 [Homo sapiens] |
| | | |
| | Putative function | Translation elongation factor |
| 35 | | |
| | Confirmation by RNAi | Not done due to failure of PCR procedure |

CATEGORY 2: FAILURE TO ENTER M-PHASE

Example 15 (Category 2)

| | | |
|----|--|--|
| | Line ID | 1216/12 |
| 5 | Category | Meiotic defects in testis: no division (no meiosis) |
| | Reversion | NR |
| | Map Position | 82F1-2 |
| 10 | Rescue ID | 2I5X-1 |
| | Rescue Sequence 1 | |
| | AAACCAAGCAACAGAAATATCTCCAGTAGAGAGCGCCACTGGAAGATCGGAA | |
| | TTTTTAGTGCTCTGCTCTGACTAACAGGTTTTAGTAGTAGTGCTTACTTTTCTAC | |
| | TACGATTTTTGTGCGGGCTAACAATTCTGTTTTCCCACTCCCTCTCTCAGTTTTT | |
| 15 | GCATGGTAACTTTTCGGTCATTGTACTGTTGTTGTTGTCTTGACACCGCAAGA | |
| | GAACAACAACAATCGGAGAAACACTGATAGCGCGGTACAGTGGGGCAGGCCA | |
| | AACTAGAACCTATACATTTAAGATGTCTCCAATTTGTGATTTTGCCTTTCAAGC | |
| | ATACTAGTTCATAGTTGATTGTTTTGTTATGTTTTGTCTTGAATGCGATGTTTCA | |
| | AGAAATCTTATTTTCGAATTACGATATTATTCTTATTCCTTTGACTTATTAAAA | |
| 20 | TAAATGAAAACGGCGAGTAGAGCAAAAGAGCGACCACTGTGGCTCCACAAGC | |
| | TCGTTTCTCTGTTTCTCATTGCGGCCAGCTCCAATTTGCGCTTATTCACACACA | |
| | CACCTCACTGCTTGCGACTGCAAATTTGTGCAGCTGAACTTTG | |
| | Rescue ID | 2I5E-1 |
| 25 | Rescue Sequence 2 | |
| | CTTGGTTTATCACCCCTCTCTCTCTCTATCGCGCGCGCGCGCTCTTTGTGGAA | |
| | ACAGGTATAACTGTTTGGCGTGAGGGAGCACGAAACTCCAGTGGAGACTTCTC | |
| | CGCATCGCCAGCGAAACAAACGATCAAAATGAATACTCTGATAACGTGTGAA | |
| | GGTGAGCAACAAAATAAAGTATAAGAAAATACCGCCACGAAAACAACAACA | |
| 30 | ATAGAAATGTCGACGCACCCTTTTCTTTTTCTCGCAAAGAACGAGGAAATGGA | |
| | GAAGCGCAAAACCACATCCCGCTTAAAGAGTCCCTTTCCCCCGCTGGAAGTGG | |
| | AAGGAAAGGCAGCTTAAAGAGGAGCGGGTGGCTTCCAGTATGTGGAAAACAA | |
| | AGCAGACGCCATTGGAATGCCGTCGTTTTTTGTTGTTGCTAAGCCGGACATGG | |
| | CAATTGTTGCTTTTGTTCGAGAGGGGGTGGTGAAACTCATAAATATCAGCT | |
| 35 | ATGGCGAGGGGGTGGGGGCAGTCTTTGTCTGACGTACCGTACTTTTAATTTCTT | |
| | GTCGCCCCGGTTTAATCCAATTTATCCAGCTTTGAATTTGCGCGG | |
| | Genomic hit, Accession No. AC007532 | |
| 40 | Annotated <i>Drosophila</i> genome genomic segment AE003603 | |
| | Annotated <i>Drosophila</i> genome Complete gene candidate CG1116 - novel | |
| | Human homologue of Complete gene candidate 2495728 HYPOTHETICAL | |
| | PROTEIN KIAA0258(aa) | |

Putative function No homologies which indicate function

Confirmation by RNAi Slight loss of G1 peak

Example 16 (Category 2)

Line ID 1344/15
Category Mitotic defects in brain: no mitosis
5 Reversion NR
Map Position 83C

Rescue ID 2F6E

Rescue Sequence

10 AGCGGGAGTGAGCCGAAAGAGAGTAATTTTGGCCGTCACCAAAAAAGTGGCT
GCATAGTGCCAAACCAATGTATGGCCGTTACGCATCTTGTTATTCTAGTGTCTT
TGGCTGTAATCAGTTTGCAGTGACAGAGGAGTTCAGTTTCAGTTGACTCGGCT
TGGTTCAGGGTTTCTGATTGCCGTCCTCTTCTCCCTCTTCGCCTACAAGTCCGC
TGTTTCGGCACCGTGACGTCACCTAGACTTACACCCCTAATCAAAGATCCACTA
15 GTTTAGATTTTCCTGCATCAACGCCATATTAAC TTTATAAGCAGTCGTTATATCT
CAAGTAGGCAAAAAAGTGTAATAGATATGTATCTAAATTGTCGTACATTCTAT
TTATTTAAATTCGTTTTTACATCCAACAGGTGTTATTTTTGAAGTCTTAGATAA
CAAACAATATTCGAATTATGTGGTAGAATACTTAGCAATATACGCACATACAT
ATACATATGAACATTATATCCAATGCTTTAAAACCGGAATATCAAGACAACAT
20 AATGCAACATCTGGTCCGAGCTATCCAGGCAATCACATTTTTTGAAGTTCCCCC
GGTTATCACACATATATCGATCATACCCCGAAATGTGTAACACAGATACAGCT
CACCATCCCTCTGATAAGATCTTATCAAGTTCGGGCTTGCTCGCTATCGTGAAT
TGGGTTGAAGGGTCCGCGATAATTGCATTGGGCATGCCATTGGTAATCACAAT
TGGCTGATAATGCTGCTGCTGCAATTCCACGGGTATGAA
25 TTCATCAATTGGTTA

Annotated *Drosophila* genome genomic segment AE003602
Annotated *Drosophila* genome Complete gene candidate CG1347 - novel protein with
myosin homology
30 Human homologue of Complete gene candidate 1503990 [dbj|BAA13194|
(D86958) KIAA0203 similar
to mouse CC1.(aa)

35 Putative function similar to coiled coil protein with ubiquitin like domain

Confirmation by RNAi Marked reduction of G1 and G2/M indicating fewer cycling
cells

40

Example 17 (Category 2)

Line ID 703/16
Category Meiotic defects in testis: segregation defects, meiotic failure (Mf-07/75)

5 Reversion R
Map Position 83B

Rescue ID 2E7E

Rescue Sequence

10 AAGCAGCCCAACAGCTACGCAAAAAGTTACTTATATTCGCAGCAAAACAGAT
TTTTTTGTTTTAATCGTAAGTATAGGAGTGAAAAATAGCGCTAGAGTAGACCT
AAGTACACAGAAAGACAAATAGGGCGAGTAAAATCGCGGTCCTGGTCATTTC
TCTGGCCTTGACCAATCCTTTGTCTGCGCTTTCGTTGGAAAAGGGGTTATGTAC
GAACTGCGTGCGTACCTAAGGCCAGATTAGTCATCGGGCAGTCATATATTCAT
15 GCAAAAAATCATTTGGTGGCCGTCGGCCTTTGTTTCGACTGTACCTTGCTCATT
TTTAATAAGCGCGACAGCAATATACACACTTTGAACCCCCATCCCACATTTTTT
CTCACCGTTTCCCCCTAATTTTCGTTTTCCCTGTGCCCATCATTCCGCTTTCGCC
ATGTCAGTGTATCGCTTCAAAATGGCGCCGAACCACATGTCTTCGTTCTCGGC
TCGTCCGCTTCGTTTCGTGCGCTCGTGTGTCGTCTCATTCGCTCTCCGAATTTTCG
20 TTTAACAAAGTGGTGCGAGCAGAGGGGCCGCTGGATTTCGAGGCAAACAACAC
ATATACCTA

Genomic hit, Accession No. CSC:AC013960

25 *Drosophila* EST several including LD15903 (AA440858), GH20091 (AI389018).

Annotated *Drosophila* genome genomic segment AE003602
Annotated *Drosophila* genome Complete gene candidate CG2922 – novel

30 Human homologue of Complete gene candidate 286001 dbj|BAA02795| (D13630)
KIAA0005 [Homo sapiens] also
NP_054757.1| HSPC028 protein
[Homo sapiens] e-179

35 Putative function Weakly similar to a region of human and murine
EIF4G2 translation initiation factors; may act as a
translation initiation factor

Confirmation by RNAi Only wild type profiles observed

Example 18 (Category 2)

| | | |
|----|---|---|
| | Line ID | 741/3 |
| 5 | Category | Meiotic defects in testis: segregation defects, meiotic failure (Mf-05/31) |
| | Reversion | NR |
| | Map Position | 88D |
| | Rescue ID | H6E |
| 10 | Rescue Sequence | |
| | | GCCTGGAGCCACCTCTAGAGCCACGGCCAAAAAATTGTGTGCCAAAAAATCG |
| | | TATGGCGTTACGCATCTTGTTATTCTAGTGTCTTTGGTTCTACAAATCTGGCCA |
| | | ATGGGATGGACGGATTTTGGGGCTTTTGCGCCCCACATATGTNTCTTACAACC |
| | | CACTCGGCCCCGGCAAGTGGGTGTCAATTACGGACATCGGCAATCCGAAGACC |
| 15 | | GGAGACCCAGAGACCCTCAGACCCCAGGGCCCCATTCGATTTCGATTTCGAGTT |
| | | GCGTGGGCCCGATCTCACATTAGTCACATCGAAGGAATGAAATAAAAAGAAAA |
| | | AACATGACGGCCGAAAAGAACTTATCCATCTTCAAAGCTCTCAGAAAATACA |
| | | AAAAACTAAAAAACTTTTGACTCTTCGTCTTTCACATTTTCGAAATCACAAAAT |
| 20 | | GTCTGCCATAAATTCCAAAGTGAACAATTGAAATAAATTTTTGCGCCATGAAC |
| | | ACGCCGACTG |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003705 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG12600 - novel protein |
| 25 | Human homologue of Complete gene candidate | CG12600- 5e-27 4240227 dbj BAA74892.1 (AB020676) KIAA0869 protein [Homo sapiens] |
| 30 | Putative function | putative cytoskeletal structural protein |
| | Confirmation by RNAi | Reduction of G1 and G2/M peaks indicating fewer cycling cells |

Example 19 (Category 2)

| | | |
|----|---|--|
| | Line ID | 773/1 |
| 5 | Category | Meiotic defects in testis: cytokinesis defects, meiotic failure (Mf-02/15) |
| | Reversion | R? |
| | Map Position | 83F |
| | Rescue ID | 2D9P |
| 10 | Rescue Sequence | |
| | | CCACCGCCCATGCCGCCATTTATTGAAAGGCCTGTACGCAGTTTGTTTTTGTTT |
| | | TTCTCTTTTTTGCTAGCTCAAACACAAAATTACTTTTTGTGGCTTGACTGGTGA |
| | | GGTCTCTCTATCTCGCTTTTTTCGTCTTTACCTCGCTCTCATTCCCTCTCTATCTG |
| | | CCCTGCTTCCTCTCACTATCTATCTACAACCTGAGGTCAACAAAATAAGTGCGT |
| 15 | | AGTCAAAAATGTAATTGAATTGATTGACAAACACAGCGAACGTAAATTTCCGT |
| | | AATGTTTAACCTTGAATTCAAATGAACAACCTGTATAAATATAATACACGGGT |
| | | AAACTCCATTTCAAAGCAAGCTAAAACATTTTAAATACATTTTAGGGAAACGG |
| | | CCAATTAAAAGAATAATATTGTGGGGATCAATCTGGGGAAAAATGCAGTATC |
| | | AGTAATGCTGAATATTTATTTTACTAAATTACAATGAAATGTCTCAAACAAAT |
| 20 | | GGGTTAATCATTTTCTTGCTCCATCTGCTTTTCCCAACTGTATCCAAGTACAAC |
| | | TACAGCATTATCCTCAACTG |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003675 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG10272 - novel protein |
| 25 | Human homologue of Complete gene candidate | |
| | | CG10272 - 2995577 |
| | | AC004490 (AC004490) |
| | | R29381_1(aa) protein includes |
| | | HMG-I and HMG-Y DNA- |
| 30 | | binding domain (A+T-hook) |
| | | found in HMG non-histone |
| | | components in chromatin |
| | Putative function | Chromosomal protein |
| 35 | Confirmation by RNAi | Loss of G1 peak indicating arrest in G2/M |

CATEGORY 3: METAPHASE ARREST

Example 20 (Category 3)

5
Line ID 1067/13
Category Mitotic defects in brain: prometaphase arrest
(overcondensation, polyploidy, scattered chromosomes with
bipolar spindle)
10 Reversion NR
Map Position 69C4-10

Rescue ID 2F8E
Rescue Sequence
15 GTTTGGGCACAGGGTTGTATTTTCATTTATTTTTGGGGGGAGTCGATACGCTCTC
TTGGCGTGGTCGAACGGTCACACTGGCCGAGAGATAACGGAAAATGTTTCAA
AGGTAAGTAAAGATTATAAACGTATTAAGCTTAATACTATAATTAGCTTACTA
TTCCAAGTATGTATAATTATTACACGTTTAAAAGGCATAACGTTAAGTGTAAC
CAAATTATATCAATGGATTTTGAATACCAATATTATTTATTTTATATTTTGAGC
20 TTAATATATTAAATCACATATATTTAAGCCTCTTTATATATGTAAATATTTTAA
TTTTATTAAAATAAATTATATATTGTTTTGTAATATGATCGAGGGCTGCCACCT
TGTGATAAATGCTTACCAACACTTTTAGGTACGCCGTTTAGTGTACGTAAGTTG
CGTACCTAGATATCCAGCGAAATCAAAACATTGAGTAAATCGTGGAATAATGG
ATGAAAATAGCTTAATCTACGGACTCGAACTGCAGGCGCGGGCTTTAACACCT
25 CAGTACGGAGAGAGCAACGATGTGTGCTTCTTCATAGCCACCAACTCCTTGAA
GCCACCAATCAGGTTCACTTAATCCAGTACGAAGA

Genomic hit, Accession No. CSC:AC020333

30 **Associated ORF**
Genscan: ORF1 predicted sequences: >16:51:11|GENSCAN_predicted_peptide_2|178_aa
MAQNISPEQSGGAGGGGSKHSDDSMVKNHAVSKRLHKELMNLMMANERGIS
AFPDGENIFKWVGTIAGPRNTVYSGQTYRLSLDFPNSYPYAAPVVKFLTSCFHPNV
DLQGAICLDILKDKWSALYDVRTILLSIQSLLGEPNNESPLNAQAAMMWNDQKEY
35 KKYLDIFYEKHKDT

>16:51:11|GENSCAN_predicted_CDS_2|537_bp
atggcgcagaatatcagccccgagcaaagtggaggcaggcggcgaggcagcaagcacagcgatgactccatgcccgtg
aaagacaatcacgccgtgagcaaaagactgcacaaggaactgatgaacctgatgatggccaacgagaggggcatctcagcgtt
40 tccggacggcgagaacatcttcaagtgggtgggcaccatagcgggtccacggaacacgggtgtatcggggcaaacgtatcggtt
gtcactggatttcccaattcctatccgtatgcagcaccctgggtgaagttcctgacgtcctgcttccatcccaatgttgatctgcagg
gcgcatctgttggacatactgaaggacaaatggcggccctgtacgatgtgcgcaccattctgctgtccatacaatccctgctgg
gcgaaccgaacaacgagagtccactgaatgcgcaggccgcgatgatgtggaatgac

| | | |
|----|--|--|
| | Drosophila Gene Hit | TBLASTX with ORF1: poor homology to several sequences including homolog of RAD6 (DHR6) (M63792), bendless (L20126) and Ubc D1 mRNA for ubiquitin-conjugating enzyme (X62575). |
| 5 | Human Homologue | TBLASTX with ORF1: ubiquitin carrier protein E2-C (UBCH10) (NM_007019.1) and ubiquitin-conjugating enzyme E2B (RAD6 homolog) (NM_003337.1). |
| | Annotated Drosophila genome genomic segment | AE003541 |
| 10 | Annotated Drosophila genome Complete gene candidate | CG10682 – vihar ubiquitin-conjugating enzyme |
| | Human homologue of Complete gene candidate | gi5902146 0B6F58A1F0665D9A ref NP_008950.1 ubiquitin carrier protein E2-C [Homo sapiens] (2.50E-50) |
| 15 | | |
| 20 | Putative function | Cyclin specific ubiquitin conjugating enzyme |
| | Confirmation by RNAi | Complete loss of G1 and G2/M peaks indicating fewer cycling cells. Immunostaining shows metaphase arrest with condensed chromosomes |
| 25 | | |

Line ID 1105/1
Category Male sterile, Female sterile, Mitotic defects in brain: prometaphase arrest
 (Overcondensation, polyploidy, fewer anaphases, high mitotic index, scattered chromosomes with bipolar spindle)
Reversion R
Map Position 69C

Rescue ID A5B

10 **Rescue Sequence**

GTACATATAATCACAATTGAGAATCGAAAACCCGACCGCCACGAAGCGCGCT
 AAATTACACGCACATACTGAAAGCCAAACAGCGGATAGCACTAGCATCCTAC
 ATATATAGACGTAGATATATAGTCATGGCGCAGAATATCAGCCCCGAGCAAA
 GTGGTGGAGCAGGCGGCGGCGGCAGCAAGCACAGCGATGACTCCATGCCCCGT
 15 GAAAGACAATCACGCCGTGGAGCAAAAGGTGAGTATCACATGGTGCAGCCTA
 AGATAATCCGCCAATATACACACACACTCACACTCACCCACAGACTGCACAA
 GGGAAGTGAACCTGAATGAATGGGCCCACCGAAAAAAGGGG

Rescue ID A5E

20 **Rescue Sequence 2**

ATATGTACTGTATAGTGGAAATTTAGTTTGATCGGTGCGGAATACGCGTCTGTT
 GCTTTTTTCAGATATTTTTTTTTTTCACTTTTGTGTGAAAACAAAATGGAAGGAGA
 ACGAGAAGAACTGTGTTTGGGCACAGGGTTGTATTTTCAATTTATTTTTTGGGGG
 GAGTCGATACGCTCTCTTGGCGTGGTCGAACGGTCACACTGGCCGAGAGATAA
 25 CGGAAAATGTTTCAAAGGTAAGTAAAGATTATAAACGTATTAAGCTTAATACT
 ATAATTAGCTTACTATTCCAAGTATGTTATAATTATTACACGTTTTTAAAAGGCA
 TAACCGTTAAGTTGTTAACCCTAAATTATATCAATGGATTTTGAATACCAATATT
 ATTTATTTTATATTTTGGAGCTTAATATATTAAATCCACATATATTTAACCCCCCT
 TTATATATGTTAAATATTTTAATTTTATTAAATAAATTATATATTGTTTGGTTA
 30 AAA

Genomic hit, Accession No. AC007328 69B-69C

Associated ORF

35 Genscan: ORF1 predicted sequences
 >/tmp/aaaaanjda|GENSCAN_predicted_peptide_1|357_aa
 MGKKAKHKKKGKGPEKTAMKADKKQAARQKKMLEKLGEANIADIQLLEAKEG
 KIEAISESVCPPTPRSNFTLVCHPEKEELIMFGGELYTGKTTVYNDLFFYNTKTV
 EWRQLKSPSGPTPRSGHQMVAVASNGGELWFPNFACISRNQSWFVFHNCRLKAA
 40 SREKVLLNFNGTVLHPANNIIVHVKLFFKKANGFKPWLLDVKLDACRFVRTNFHPF
 VRIIFDLFKDFSTINHTCPYVVLRSRMRYIVRRSPRLVHPIVDVPAIGHTRPRRKA
 AVRGIGCAHRCPLIRMATPCRTNVMMTMLMRGSVRSRVMAICCYRRPAIAIARRRHP
 TAIHSQEVAERLGGLLYPDIQRTNP

45 >/tmp/aaaaanjda|GENSCAN_predicted_CDS_1|1074_bp
 atggggcaaaaaggccaaacacaagaagaaggggcaaaaggggcccgagaaaacggccatgaaagcggacaaaaagcaggcgg
 cgcggcaaaaagaaatgctggaaaaactgggagaagcaatatagctgatatcatccaattgctggaggccaaggaggggcaag
 attgaagccatcagtgaatccgttgcccgccaccaactccacgatccaatttcaccttagtttgccatccggaaaaggaggagctc

atcatgtttggcggcgaactgtacactggcacaaaaaccacagtgtataacgatttggtcttttacaacacaaaaaccgtcgagtgg
aggcagctgaaatcgccatcgggacccacgcccagaagtggacaccaaattggtggctgtggccagcaatggaggagaactct
ggtttccgaacttcgctgtataagtcgcaatcaatcctggtttgtgtccacaattgtcgtctgaaggcggccagtcgtgagaaggt
cttactcaactttaatggaacggttctacatccggccaataacataatagttcacgtcaagctgtttaaaaaggccaacggtttaage
5 cttggttattagacgtaaaactcgatgcttgtcgtttgtgcggaccaacttccatccgtttgtacgcattatattcgatctcttcaaagat
ttctccaccataaaccacacgtgcccataatgiggctcctccgatcgatgcggtatattgtccgcatccccacgacttgtgcacccc
atcgtagatgttccggctattgggcacactcgccctcgacggaaggccgcttcgtggcataggggtgtgctcatcgctgccctct
gattcggtatggcgactccgtgtcgtaccaacgtggtgatgatgacgctgatgaggggctcgggtgagatcgagggtgatggcgatt
tgctgctaccgccgacccgccattgccatagccgctcggcgccaccccactgccattgccactcccaagaagttgctgaacgc
10 ctcggtggtcttcttaccggacattcagagaaccaatccgtag

***Drosophila* EST** several ESTs including LD04777 (AA201675)

All other entries as for 1067/13.

Example 21 (Category 3)

| | | |
|----|--|---|
| | Line ID | 1407/13 |
| | Category | Mitotic defects in brain: (weak overcondensation, metaphase with bipolar spindle) |
| 5 | Reversion | NR |
| | Map Position | 92B1-3 |
| | Rescue ID | 2D3P |
| 10 | Rescue Sequence 1 | ATCACGAATTTGACATTGCTACCACATTCGGTGCGTGGACTCTGAAAGCTCTG AGTGTTTTGTTTATGCAAAGCTTTTTTGGACTATCGCGTGGTAAGTAGCCGAAA GAGAAAGCTCTCTTATACGGAAGATGAAGAGTGTGATTCATGAAAATGTATA AGAACGCGGGTCCAAAAAGTCAAGGGAGTTCTAGTGAAATGAAAAGTTCCAA 15 AGGTTTTGAAATCGTTTTATTTTCTCGTTCGTATAATTATTGGGTGTCGATCTTT GTTGGGCAGTGTAAGCACAACTTTGAGCTTCATCATAATCATATGTAA AGCCGGGACGAAAGCTTATGATTCTGTAAAGTGTCCGCCCAAGATAACATTTT TCCAGCCCTTCAAATCTTCAAATAAATACGGCTTAAGGCGAGCAAATTTGTAA ATCAAATGATTTGTTAAATAAACATTATATGTATTTTATCATGCCAGGTTAGAA 20 CACATTGTGCTGATGCAAATAAAATTCCAATTAAACGCCCTGAATGGGAAGA TGACGCATCTTTAATGGGAATATTATGGTAAATTTAATA |
| | Rescue ID | 2D3E |
| | Rescue Sequence 2 | TNCGTGATTATCAGCGTTAATTGTACAATATTATGATTTATTTCGAGCTGTAAAT 25 CTTACAGCAAGCACAACTGTAATTATACCACTTAGAATTCCGCGGAATTAA TTCTTGAAGACGAAAGGGCCTCGTGATACGCCTATTTTTATAGGTTAATGTCAT GATAATAATGGTTTCTTAGACGTCAGGTGGCACTTTTCGGGGAAATGTGCGCG GAACCCCTATTTGTTTATTTTCTAAATACATTCAAATATGTATCCGCTCATGA 30 GACAATAACCCTGATAAATGCTTCAATAATATTGAAAAAGGAAGAGTATGAG TATTCAACATTTCCGTGTCGCCCTTATCCCTTTTTTGCGGCATTTTGCCTTCCT GTTTTGCTCACCCAGAAACGCTGGTGAAAGTAAAAGATGCTGAAGATCAGTTG GGTGCACGAGTGGGTACATCGAACTGGATCTCAACAG |
| 35 | Drosophila EST | LD05707 (AA246767) |
| | Annotated Drosophila genome genomic segment | AE003727 |
| 40 | Annotated Drosophila genome Complete gene candidate | CG7444 - very short ORF with EF hand homology |
| | Human homologue of Complete gene candidate | none |
| 45 | Putative function | Possible calcium binding protein |

Confirmation by RNAi Slight loss of G1 peak

Example 22 (Category 3)

| | | |
|----|---|---|
| | Line ID | 1439/7 |
| 5 | Category | Mitotic defects in brain: prometaphase arrest. (overcondensation, polyploid, no anaphases, scattered chromosomes with bipolar spindles) |
| | Reversion | ? |
| | Map Position | 96F10-14 |
| 10 | Rescue ID | G3X |
| | Rescue Sequence | |
| 15 | | GTCGGATGTAGAAGACGTGCCCCGAAACCCAGTTAGAAATCGATGTCAGCGAT GGCGCCGGACTGGAGGATGAGGATGATGACGATATGGAACAGATTACAGCTC AGAAGGTAAGGTAAATCGTAACAGAGCTTTTAAATACGCAAGTAATCACATTC TGATATCCCTAGGTTCTGGAAATCATAGAAACCGCGTGGATAAATGAAATGTG TGCGCCGGAGATCCTGCCCAGCCAGACGGACATGCTGGAGCTGATGGTCTCCC AGGTGGCCCATATGGAGGAGCAGATGCGCGATCTGGACAAGAACGATTTCG AGCGGTGGTGCACTCCATGGAACCTGGAGAGGGTGCGCTACATAATGGCCAGT TATCTGCGTTGCCGCCTGCAGAAGATCGAAACCTTCACGCAGCACATCCTCAA 20 CCAGGAGGAGAGCCGTGAGCCGGATGACAAACGTCTGTCTCCCGAGGAGACT AAGTTCGCCCAGGAGTTTGCCAGTAAT |
| | Genomic hit, Accession No. AC007825 | |
| 25 | Annotated <i>Drosophila</i> genome genomic segment | AE003754 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG14549 – novel |
| | Human homologue of Complete gene candidate | none |
| 30 | Putative function | no homologies which indicate function |
| | Confirmation by RNAi | Only wild type profile observed |

Example 23 (Category 3)

Line ID 1466/4
Category Mitotic defects in brain: metaphase arrest.
 (overcondensation, no polyploidy, fewer anaphases, metaphase
 with bipolar spindle)
Reversion NR
Map Position 72F

Rescue ID E5E

10 **Rescue Sequence 1**

GGCTGGATGCGATTCGCTTTCGGATTCGGATGGATTCAGCCGCTGTCTCGACA
 CCGCCGCAACCGCTCTCGGGAGTTTGAAAATTTGAAATGAGCGGATTCGCGTT
 GCGAAGGCGAGCTAGCGTTGCAGGCAGTGTGGCCAGATGCCGCGTGCGAACG
 TATTCTCGAATGCAATCGGCCGAGTGCAGATGCACTAAAAATAACCCACTTCC
 15 AGTGACTGGAAATTAAGATCAAGGNAATAGATTTTATAAAAACTTATATGAGT
 AAAAATTTTAAAATTGTGGAGTCAACCTAAATTATAAGCAACTAATTTATAAC
 ACAAGTAAAGAATGATATTAAGTAACTTTTAAATAATATTCCATTATGCTTA
 CGCTCAATTTATGAACAAATGTTTTCTCGATCCTTAGGTAAAGTTTCGAGTTTC
 GCGACTANATTTATTAATAAATTAAGAACATCTCCATTTATGTACACATTAAAG
 20 ATTTATGAGCGGTAATATTAGCTGGTTGAC

Rescue ID E5P

Rescue Sequence 2

ATCCAGCCAAGATATCCTATCGTGCAGCTGAAACCCGAAACCCGAATCCGAGT
 25 TCGAAACGAAACGAATCGCAGTGGTGGTTTCTCTCTCGCTCTCTAGCTCTCCCT
 CTCTCTCGCGTGTGTGTATGTGTGCGAGTGGCAGGAAAAGTGCGAAGCCGAAA
 TCTTTTCTAGCTGAAAGAAAGCGCAACTTCAATTAGCGAAAAGCAAGAGTAGCT
 AACAAAAAGAAAAGCGGATCGAAAAGTAAAGAAAAACAAAAAACA
 AAAGCAACAAATCGAAATGGCAAGCGAAGTGGCCCAAATACCCGCCGAGGG
 30 AAACGCCCCGAGTGGCGGCGGCGGAAAAATCAGAGGAGCCGGAAAAGTCAG
 CGGCCCCCGCCAGCGGACTCAGCGGCCGCTCCAGCTGCCGCCCCCGCAGTGGA
 GAAGGCTGAGGATGCCGATGGCGAAAAAAGGACGGCGAGGCCGGAAAGCA
 GGACAAGCAGCAGGATGGC

35 **Genomic hit, Accession No.** CSC:AC020154

Associated ORF

Genscan ORF: ORF2 predicted sequences

>21:06:03|GENSCAN_predicted_peptide_5|415_aa

40 MASEVAQIPAEETPAVAAAEKSEEPEKSAAPPADSAAAPAAAPAVEKAEDADGE
 KKDGEAGKQDKQDGEPPKKDEAVAAPVATKSEAPPAQKFNVHKTNFEKDIIYL
 YQFSRTPLLPSLSPYCLKVETWLRLVGLKYENVDHKMRFRSKKGQLPFIELNGEEI
 ADSAIIKELSSKYEKYLD SGLTAEQRNVSYATIAMLENHLIWIIFYWRAKYPDNV
 LKGYKVN LQHALGLRLPNSILNFFFKITFGRKGTKKLKAHGIGVHSAEEIEEF GKD
 45 DLKVLSEMLDCKPFFFGDEPTTLDVVAFAVLSQLHYLSKDIA YPLRDYMTEKCPN
 LIGHVSRMKDKCFPDWDEICTKLDLNAHIPKPEPETKEGKEGGEQEKSNEQEGTE

GDKIEKELEKDKSNEKESTEENKEKEETK

>21:06:03|GENSCAN_predicted_CDS_5|1248_bp
atggcaagcgaagtggcccaaatacccgccgaggaaacgcccgcagtggcggcgggcggaataatcagaggagccggaaaa
5 gtcagcggccccgccagcggactcagcggccgctccagctgccgccccgcagtggagaaggctgaggatgccgatggcga
gaagaaggacggcgaggccggaaagcaggacaagcagcaggatggcgaggagccccaaaaaggacgaggcgggtggcagc
accctggcgaccaaatcggaagccccgcccgcagaaattcaatgtgcacaagaccaacttcgagaaggacatcatctatct
gtaccagttctcgcgcacccactgctgccctccctgtcgcctactgcctgaaggtggagacctggctgcgtcttgtgggcctga
aatacgagaatgtcgatcataagatgcgtttccgctccaagaagggtcagctgccgttcatcgagctgaatggggaggaaatcgc
10 cgattcggccatcatcatcaaggaactgtcgtccaaatacgagaagtacctggactcgggactcaccgccgagcaaaggaatgt
ctcgtacgccacgattgccatgctggagaacctctcatctggatcatcttctactggcgcccaagtatccggacaatgtgctcaa
gggctacaaggtcaactgcagcacgccctcggcctgcggctgcccaactcgattctgaacttcttcttaagatcaccttggctgc
aagggcacgaagaagctgaaggcgcacggcatcgggtgtccacagcgccgaggagatcgaggagttcggcaaggacgacctg
aaggtgctcagcgagatgctcgaactgcaagccttcttcttcggcgacgagcccaccacctggatgtggtggccttcgctgtcct
15 ctcgcagctccactatctgtccaaggacattgcgtatccgctgcgcgactacatgaccgagaagtgccccaaacttgattggccacg
tatctcgcattgaaggacaagtgttccccgactgggacgagatctgcacgaagctggacctcaatgcgcacattcccaagccag
agcccgagaccaaggaggggcaaggagggtggcgagcaggagaaatcaaacgaacaggaggggcactgagggcgacaagat
cgagaaggagttggagaaggacaagtcaaacgagaaggagtcgaccgaggagaacaaagagaaggaggaaacaaagtaa

Drosophila Gene Hit rescue sequence and TBLASTN with ORF2: failed axon connections (U21685)
Human Homologue BLASTX with fax: Metaxin 1 and 2 (Q13505 and AF053551)
Drosophila EST several including LD31362 (AA951078 similar by BLASTN to U21685 failed axon connections)

Annotated Drosophila genome genomic segment AE003527
Annotated Drosophila genome Complete gene candidate CG4609 – fax failed axon connections
connections
Human homologue of Complete gene candidate 4505281
ref|NP_002446.1|pMTX|
metaxin>gi|3024205|sp|Q13505|MTXN_HUMAN
METAXIN (4e-06)

Putative function Drosophila fax is a dominant genetic enhancer of the Abl mutant, developmentally expressed in axons of the CNS

Confirmation by RNAi Weak reduction of G1 and G2/M peaks indicating fewer cycling cells

| | | |
|----|--------------------------------------|---|
| | Line ID | 262/20 |
| | Category | Mitotic defects in brain: metaphase arrest. (overcondensation, polyploidy, aneuploidy, few anaphases, high mitotic index, metaphase with bent bipolar spindle) |
| 5 | Reversion | NR |
| | Map Position | 72F |
| | Rescue ID | G6E |
| | Rescue Sequence | |
| 10 | | AGCTGCACGATAGGATATCTTGGCTGGATGCGATTTCGCTTTCGGATTTCGGATG GATTCAGGAGCCGCTGTCTCGACACCGCCGCAACCGCTCTCGGGAGTTTGAAA ATTTGAAATGAGCGGATTTCGCGTTGCGAAGGCGAGCTAGCGTTGCAGGCAGT GTGGCCAGATGCCGCGTGCGAACGTATTCTCGAATGCAATCGGCCGAGTGCA GATGCACTAAAAATAACCCACTTCCAGTGACTGGAAATTAAGATCAAGGAAT |
| 15 | | AGATTTTATAAAAACTTATATGAGTAAAAATTTTAAAATTGTGGAGTCAACCT AAATTATAAGCAACTAATTTATAACACAAGTAAAGAATGATATTAAGTAACTT TTTAAATAATATTCCATTATGCTTACGCTCAATTTATGAACAAATGTTTTCTCG ATCCTTAGGTAAAGTTTCGAGTTTCGCGACTAGATTTATTAAAATTAAGAACA TCTCCATTATGTTCCC |
| 20 | Drosophila EST | several including LD28084 (AA949260) |
| | All other results as for line 1466/4 | |
| 25 | | |

Line ID 262/22
Category Mitotic defects in brain: metaphase arrest.
(overcondensation, polyploidy, few anaphases, high mitotic index,
metaphase with bent bipolar spindle)
5 **Reversion** NR
Map Position 72F

Rescue ID F1E

Rescue Sequence 1

10 AGCTGCACGATAGGATATCTTGGCTGGATGCGATTTCGCTTTCGGATTTCGGATG
GATTCAGGAGCCGCTGTCTCGACACCGCCGCAACCGCTCTCGGGAGTTTGAAA
ATTTGAAATGAGCGGATTTCGCGTTGCGAAGGCNAGCTAGCGTTGCAGGCAGT
GTGGCCAGATGCCGCGTGCGAACGTATTCTCGAATGCAATCGGCCGAGTGCA
GATGCACTAAAAATAACCCACTTCCAGTGACTGGAAATTAAGATCAAGGAAT
15 AGATTTTATAAAAACTTATATGAGTAAAAATTTTAAAATTGTGGAGTCAACCT
AAATTATAAGCAACTAATTTATAACACAAGTAAAGAATGATATTAAGTAACTT
TTTAAATAATATTCCATTATGCTTACGCTCAATTTATGAACAAATGTTTTCTCG
ATCCTTAGGTAAAGTTTCGAGTTTCGCGACTAGATTTATTAAAATTAAGAACA
TCTCCATTTATG

20 **Rescue ID** F1P
Rescue Sequence 2

GTGCAGCTGAAACCCGAAACCCGAATCCGAGTTCGAAACGAAACGAATCGCA
GTGGTGGTTTCTCTCTCGCTCTCTAGCTCTCCCTCTCTCTCGCGTGTGTGTATGT
25 GTGCGAGTGGCAGGAAAAGTGCGAAGCCGAAATCTTTTTAGCTGAAAGAAAG
CGCAACTTCAATTAGCGAAAAGCAAGAGTAGCTAACAAAAAGAAAAGCGGAT
CGAAAAGTAGAGAAAAACGAAAAAAAAAAAAACCAAAGCAACAAATCGAAATG
GCAAGCGAAGTGGCCCAAATACCCGCCGATGAAACGCCCGCAGTGGCGGCGG
CGGGAAAAATCAGAAGAGCCGGAAAATCAGCGGGCCCGCCAGCGGGACTCTG
30 CGGGCGCTCCAGCTGCCGCCCCCGCAGTGGAGAAGGCTGAGGATGCCGATGG
CGAA

Drosophila EST several including LD28084 (AA949260), LD38479 (AI518768)

35 Other results as for line 1466/4

| | | |
|----|------------------------|--|
| | Line ID | 262/3 |
| | Category | Mitotic defects in brain: Metaphase arrest (overcondensation, polyploidy, aneuploidy, no anaphases, high mitotic index, metaphase with bipolar spindle) |
| 5 | Reversion | NR |
| | Map Position | 72F |
| | Rescue ID | H3E |
| | Rescue Sequence | |
| 10 | | AGCTGCACGATAGGATATCTTGGCTGGATGCGATTTCGCTTTCGGATTTCGGATG GATTCAGGAGCCGCTGTCTCGACACCGCCGCAACCGCTCTCGGGAGTTTGAAA ATTTGAAATGAGCGGATTTCGCGTTGCGAAGGCGAGCTATCGTTGCAGGCAGTG TGGCCAGATGCCGCGTGCGAACGTATTCTCGAATGCAATCGGCCGAGTGCAG ATGCACTAAAAATAACCCACTTCCAGTGACTGGAAATTAAGATCAAGGAATA |
| 15 | | GATTTTATAAAAACTTATATGAGTAAAAATTTTAAAAATTGTGGAGTCAACCTA AATTATAAGCAACTAATTTATAACACAAGTAAAGAATGATATTAAGTAACTTT TTAAATAATATTCCATTATGCTTACGCTCAATTTATGAACAAATGTTTTCTCGA TCCTTAGGTTAAGTTTCGAGTTTCGCGACTAGATTTATTAAAATTAAGAACATC TCCCTTTATGTTC |
| 20 | | Other results as for line 1466/4 |

Example 24 (Category 3)

| | | |
|----|-----------------|---|
| 5 | Line ID | 238/20 |
| | Category | Mitotic defects in brain: metaphase arrest (overcondensation, metaphase with bipolar spindle) |
| | Reversion | NR |
| | Map Position | 75E1-3 |
| 10 | Rescue ID | D7E |
| | Rescue Sequence | TTCAGTCGCGCATTTCACCGTTTCCGAATCGGACGAACCGGGCGTGATTGCTC TCCTGCTGCTTTTCGAGATCGGAGTCCCGATAAGGATATAACTACAACCTAAAG AGGAATCCAAGCCTCCTCCTGCCGCTAGTTTCGAAAAGTAAATAGAGTACTTG TTATCAACTGGGGAAGCGGAGATACATAGCTCCGATATTCCTGTGAAAGCCAG 15 ACAAACGGGATACCAACGAACAATCGCCATGTGCGTCGTCGTCCTTCTCGTTT CACACATCGTGCGATAAAAATACCGCTTTGCTTTTTGTGTTTATTTAAAAATTT TGGTTAGGAAGTGAAGTTCGAACTCGTGACGTTTGCATTTTCACAACAACAAAA AGAGCAAAACATAGCAGAAGAACCCAGAAAGAACAGGAACAGAAACCGTT GACCGAGTGCCAGTGTGAAGGTCTAGGCACAAAGAACGCTACCAAGAACTCT 20 TGGGAGTTAGGGAGGCTCTTTACAATGACAACATTGCACCAAAGATGGACTCT CTCTCTAAAATGCATTTTCATACCAATATTTACTTT |

Drosophila EST several including LP04802 (AI260815)

| | | |
|----|--|--|
| 25 | Annotated <i>Drosophila</i> genome genomic segment | AE003519 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG3979 - novel gene with homology to sodium- dependent dicarboxylate transporters |
| 30 | Human homologue of Complete gene candidate | 3e-87 4506979 ref[NP_003975.1 pSLC13A2 UNKNOWN >gi 2499523 sp Q13183 NDC1 _HUMAN RENAL _SODIUM/DICARBOXY |
| 35 | | |
| 40 | Putative function | sodium/dicarboxylate transporter |
| | Confirmation by RNAi | Only WT profiles observed |

Line ID 490/9
Category Meiotic defects in testis: segregation defects, multipolar spindles (Mul-02/29)
Reversion NR
5 **Map Position** 95C1-8

Rescue ID I4E

Rescue Sequence

10 GCTCTGCCGCTTCAACCGCCCGCGTTCTGTGTGTTGGTGTGCCGCGACGTAGG
TG TAGGGTCCGCTGCACACGTGTGTGTGGGAGCGCGCGAGAGCGGGAGAAGA
GCAGAACGTTTTTGGGCGGCTAGTGGTGGCACCGTGAGCATGCCGGTCGTCGT
AAGATAGGCTTAGGAACACTCAGAGAAAATTTGTTTAGCTCAGCATTTTCCTA
TTATTGAAATCATTTATTTGATGGTCTATGGGGGTTTCTTTCGTAGTTATTCAT
AGATCGGCGATTTAAGCTACGCTTAAAGGGTAATTTGTCTGAGATATCTTTGT
15 CATTTAAAGTTAAGTCTCAGCTTATCCAAAAGTCAGTTATTGGAAAAAAGGAG
CCAGCTTTTCAGCAGAGTTCGGCTTAAGCGCTTATTATCATATTAACCAGCTTA
ATTAATGTATCTTTTAAATTGTTATATGCATTAAATCACTAATTAAGGTGATTA
CCATTTGTACGTTTTTAAATTAAAGTATTTTGATTTTCACTAATAACAGGCTCTAA
GCTGATCCAAATCTACAAGCTTAGTTTTTGAATAGTCTTCACATGTTGACTTTT
20 ATTCTCT

Genomic hit, Accession No. CSC:AC015160

Other results same as 238/20

25

| | | |
|----|-----------------------------------|--|
| | Line ID | 660/3 |
| | Category | Meiotic defects in testis: cytokinesis defects,abnormal spindles. (Ab-01/03) |
| | Reversion | R? |
| 5 | Map Position | 75E |
| | Rescue ID | H8E |
| | Rescue Sequence | |
| 10 | | GCTCTGCCGCTTCAACCGCCCGCGTTCTGTGTGTTGGTGTGCCGCGACGTAGG TGTAGGGTCCGCTGCACACGTGTGTGTGGGAGCGCGCGAGAGCGGGAGAAGA GCAGAACGTTTTTTGGGGCGGCTAGTGGTGGCACCGTGAGCATGCCGGTCGTCGT AAGATAGGCTTAGGAACACTCAGAGAAAATTTGTTTAGCTCAGCATTTTCCTA TTATTGAAATCATTTATTTGATGGTCTATGGGGGTTTCTTTCGTAGTTATTCAT AGATCGGCGATTTAAGCTACGCTTAAAGGGTAATTTGTCTGAAATATCTTTGT 15 CATTTAAAGTTAAGTCTCAGCTTATCCAAAAGTCAGTTATTGGAAAAAAGGAG CCAGCTTTTCAGCAGAGTTCGGCTTAAGCGCTTATTATCATATTAACCAGCTTA ATTAATGTATCTTTTAAATTGTTATATGCATTAAATCACTAATTAAGGTGATTA CCATTTGTTCGTTTTAAATTAAAGTATTTGAATTTC |
| 20 | Genomic hit, Accession No. | CSC:AC015160 |

Other results same as 238/20

Example 25 (Category 3)

Line ID 273/18
Category Mitotic defects in brain: metaphase arrest
 5 (overcondensation, very high mitotic index, few polyploids, metaphase with bipolar spindle)
Reversion NR
Map Position 75E

10 **Rescue ID** D1E
Rescue Sequence
 AACTGGGCTAAAACCAGCTGAAAAGTAAAATATTTGGAGAAG
 GAAAGCCTTAAGTTCCTCTCTACGCTTCGTACACGTAATGTGCGTGGTTTAATC
 TACGTTAAACAAGTGGAACCATGTTACGTGCCGTGGCTTTGTGTGTGTCAG
 15 TGGTGCTCATAGCACTATATACGCCAACTTCTGGGGAATCCAGTCAGAGCTAT
 CCCATTACCACGCTAATCAACGCGAAATGGACGCAGACGCCCCTATATCTGGA
 AATCGCCGAGTATCTGGCCGATGAGCAGGCGGGCCTCTTCTGGGATTACGTTT
 CGGGGGTGACAAAGTTGGACACGGTTCTCAACGAATATGGTTTGTGTTTATAA
 GTCATGGAGAACCCGCATTAAAGAGCTTTTTATATTCTCCTCAATGTGAATCC
 20 GAATCCATATAAAATC

Genomic hit, Accession No. AC015160

Associated ORF

Genscan: >ORF2 predicted sequences

25 >16:57:34|GENSCAN_predicted_peptide_5|1548_aa
 MLRAVALCVSVVLIALYTPTSGESSQSYPIITLINAKWTQTPLYLEIAEYLADEQA
 GLFWDYVSGVTKLDTVLNEYDTESQQYNAALELVKSHVSSPQLPLLRLVSMHS
 LTPRIQTHFQLAEELRSSGSCQSFTFAQVGSELACSFNELQKKLEVPLAKDSLDA
 VVTYSFDHIFPGSENNTRTVVLYGDLGSSQFRTYHKLLEKEANAGRIRYILRHQLA
 30 KKDKRPVRLSGYGVELHLKSTEYKSQDDAPKPEAGSTSDEDLANESDVQGFDFK
 VLKQKHPTLKRALDQLRQRLQGNDEIAQLKAWEFQDLGLQAAAAIAEIQGDET
 LQILQYTAHNFPMMLARTLLAHKVTDGLRAEVKHNTAFGRSLNVAPPDGFALFING
 LFFDADTMDLYSLIETLRSEMRVLESLSNNVRGSLASSLLALDLTASSKKEFAIDI
 RDTAVQWVNDIENDVQYRRWPSSVMDLLRPTFPGLRNIRKNVFNLVLVVDAL
 35 QPTARSVIKLSESVIHQAPIRLGLVFDARDANEDNLADYVAITCAYNVVSQKKD
 ARAALSFLTIDIYAAVGETKVVTKKDIVKQLTKEFTSLSFKAEEFLEEDSTYDYGR
 ELAAEFIQRLGFGDKGQPQALLNGVPMPSNVVTADSDFEEAIFTEIMTHTSNLQKA
 VYKGELTDNDVAIDYLMNQPHVMPRLNQRLSQEDVKYLDINGVAYKNLGNVG
 VLNRLSNRDMTATLMDNLKYFGGKKSTELIGRASLQFLTIVVFADLETQGRDLL
 40 THALDYVQSGESVRVAFIPNTESSASSRRNLNRLVWAAMQSLPPTQATEQVLK
 WLKKPKEKIEIPTQLEDILGSTELHLKMLRVYSQRVLGLNKSQRLVIGNGRLYGPL
 SSDESFDSDAFALLARFSSLQYSDKVRQVLKESAQDVNEEFNSDTLLKLYASLLPR
 QTKTRFKLPTDLKTDHSVVKLPKQENLPHFDVAAVLDPASRAAQKLTPILILLRQ
 VLNCQLNLYLIPVPQHSDMPVKNFYRYVVEPEVQFEANGGRSDGPLAKFSGLPAN
 45 PLLTQQQLQVPENWLVEAVRAVYDLNKLTDIGGPVHSEFDLEYLLLEGHCFDAA
 SGAPPRGLQLVLGTQSQPTLVDTIVMANLGYFQLKANPGAWSLRLREGKSADIYA

ISHIEGTNTHHSAGSSEVQVLITSLRSHVVKLRVSKKPGMQQAELLSDDNEQAAQS
GMWNSIASSFGGGSANQAATDEDTETINIFSVASGHL YERLLRIMMVSLKHTKSP
VKFWFLKNYLSPQFTDFLPHMASEYNFQYELVQYKWPRWLHQQTEKQRTIWGY
KILFLDVLFPLNVRKIIFVDADAIVRTDIKELYDMDLGGAPYAYTPFCDSRKEMEG
5 FRFWKQGYWRSHLMGRRYHISALYVVDLKRFRKIAAGDRLRGQYQALSQDPNS
LSNLDQDLPNNMIHQVAIKSLPDDWLWCQTWCSDSNFKTAKVIDLCNNPQTKEA
KLTAQRIVPEWKDYDAELKTLMSRIEDHENSISRDSA VDDSVDDSVDEVTTVTPS
HEPKHGEL

10 >16:57:34|GENSCAN_predicted_CDS_5|4647_bpatgttacgtgccgtggctttgtgtgtgtctgtggtgctca
tagcactatatacgccaacttctggggaatccagtcagagctatcccatcaccacgctaataacgcgaatggacgcagacgcc
cctatatctggaaatcgccgagtatctggccgatgagcaggcgggctcttctgggattacgtttcgggggtgaccaagttggaca
cggttctcaacgaatatgataccgagtcgcaacagtacaatgccgccttgagctggtcaagagccatgtgagttctcccaattg
cccctgcttaggctggtggtatccatgcatagcttgacgccccggatccagaccacttccagttggccgaggaactgaggagca
15 gtggctcttgctagagctttactttgcccaggtgggttccgaactggcctgcagctttaacgagctgcagaagaagctggaagtgc
cgctcgccaaggatagcttgatgcttctgtgtcacctacagcttgatcacatttccctggcagtgagaacaatacccgcaactgt
ggctactatacggcgatttgggaagctctcaattccgcacctatcacaaactattggaaaagggaagccaatgctggccggattcgtta
catcttgcgtcatcaattggccaagaaggacaagcgaccggtacgactttcgggctatggagtggaaactccatctgaagtcaacg
gaatacaagagtcaggatgatgctccaaagcccgaaagctggttccacttctgatgaggatttggctaataatcgacgtccagg
20 gctttgatttcaaggtgctgaagcagaagcatcctacacttaagagagcgctggatcaactgcgtcagaggcttcttcagggaac
gatgagatcgcccaattgaaagcatgggagttccaggatttgggtctccaggcggccgctgctattgcagaaatacagggtgatg
aaaccctacaaattcttcaatatactgccataatttccccatgttggccagaaccctgctggcccaaggttacggatggcttaag
ggcggaggttaaagcataatacgggaagcatttgggaagaagcttgaatgtagcgctccagatgggtgcccttttcatcaatggactctt
cttcgatgctgacacaatggatctgtatccctgattgagacgctgcgctcgagatgcgtgttctcgagagtctgcacagtaataat
25 gtgaggggaagccttggcagctccttgcctcgttgatctgacggcctccagcaaaaaagaattcgccatcgacatccgtgaca
ctgcagtacagtggtcaacgatattgaaaacgatgtgcagtaccgcaggtggccctcatcggtgatggatcttttgcgtccaacct
ttcctggcatgttaaggaataatccgaaagaatgtgttcaatttggctcctagtggtagacgcgctgcagcccacagctagaagtgttat
taaactgtcagagtcgtttgtcatccatcaagctcccattcgcttgggttgggttctgatgcgagggacgccaacgaggataatcttg
cagattacgtagccatcacgtgcgcctataactatgtgagtcagaaaaaggatggcgagctgctttaagtttctcaccgacatct
30 acgcagcagttggtgagaccaaagtggtcacgaaaaagacatagtcgaagcaactaacgaaggaatttcatcattaagctttgc
caaagcggaggagttcctggaggaagattccacgtacgactatggcagggagctcgcagcagagttcattcagcggctgggatt
cggagacaagggacaacctcaggccttgttgaatggtgttccaatgccagcaacgttgtgaccgccgatagcgacttcgagga
ggctattttcaccgagattatgaccacaccagcaatctccaaaaggctgtgtacaaaggtgaactgacagacaacgatgtagcca
ttgattatctgatgaatcaacctcacgtgatgccagattgaatcagcgaatectaagccaggaggatgtgaaatatcttgatattaac
35 ggcgtggcctacaaaaatcttggcaatgttggagttaaatcgtctgtctaaccgggatagaccgctacgctaattggataatcttaa
atactttggtggcaagaagtctacggagcttattggccgagcatccctacagttcctaacgatttgggtgtttgctgatttgaaactg
accagggctgagatctgtcaccatgccttgactatgtccaaagtggagagagtggtgcgagtcgattcattccaaacactga
aagctcttcgcctcaagccggaggaatcttaatcgattgggttgggtgccatgcagagcttccaccaactcaagccacggagc
aggttctcaagtggttaagaaaccaaaggagaaaattgagataccactcagctcgaggatatcctgggatctacagagctgca
40 cctgaagatgttgagagtttattccagcgagtggttgggtctaaataaatccagcgtttgggtcatcggtaatggcggtttatggg
ccccttctgcggatgaaagctttgatagcgccgatttgccttgcctagccaggttcagttctctacagtatagcgataaggtgcgtca
ggctctgaaggaatctgctcaagatgtcaatgaggaattcaacagcgatacattgcttaagttgtatgccagcctgcttccaggca
aaccaaaactcgctttaagctaccaacggacttaaaaaccgatcactcggttgtaaaactaccgccccaaacaggagaatcttccc
atlttgatgttgcggcgttttggatcccgcctcccgagcagctcaaaaactaacgccaatacttatttgcctcgtcaagtgtgaact
45 gccaatgaactatacctgattcccgtccccagcacagcgatatgccggtgaagaacttctacagatacgttgtgaaccggag
gtccaattcgaggcgaatggaggccgatctgatggctcttggccaaattcagtggttgcagccaatcctctgctgaccagca
gctgcaggttcccgagaactgggttggtcgaagctgtgagagcagtttacgatctggacaacattaagttgaccgatattgggtggac
ctgtgcacagcgaattcgatctggagtatctgctgttggagggtcactgctttgatgctgctagcggcgctccgcccagaggacttc

agttggtgttggtaccagagtcacctaccttgtagatactattgtgatggcgaatttgggttattccaacttaaagccaatcca
 ggagcttggtccctacgcttgctgaaggcaaatcggcggatatttatgcaatcagccacattgaagggaacaaatacccatcattc
 ggctggctcttctgaagttcaggttcttataacctccttgcatcccatgttgtaaaattaagggtgtctaagaagccaggcatgcag
 caggcggaaactcctgtcagatgacaacgaacaggcagcgaatcaggcatgtggaacagcatcgccagcagttttggcggcgg
 5 cagtgccaaaccaagcagccactgatgaggatacggaaaccatcaacattttctgtggcatcgggacactgtacgaacgtcttct
 aaggatcatgatggttcgctgctaaagcacacaaaatcacctgtgaagttctggttctgaagaactatcttcgccgcaatttacgg
 atttccttcctcacatggccagtgagtacaactccagtacgaattgggtccagtacaaatggccccgctggctgcatcagcaaacgg
 aaaaacagaggaccatttggggctacaagatccttttctggacgtgctcttcccgtgaatgtgaggaaaatcatttctggtgatgc
 cgatgccatcgtaagaacggatataaaggagttgtatgacatggacctcggaggagcacccctatgcctacacgccattctgcgatt
 10 cccgcaaagagatggagggttccgattctggaagcagggatactggcgaagccatctgatgggcaggcgttaccacatttccg
 cctgtacgtggtggacttgaagagattccgcaagattgcggcaggagataggctaagaggccaataaccaggcacttagccagg
 atccgaacagcttatccaatttgatcaggacttgcacaacaacatgatccaccaggtcgccatcaaataccctgcccagcactgg
 ctatggtgccaaacgtggtgcagcgacagcaactcaagactgctaaagtgattgatttgcacaacccgcagaccaaggagg
 ccaaaactcacggccgcccagaggattgtgcccgaatggaaggactacgatgccgagctgaagaccctgatgtctcgcacgag
 15 gatcatgagaattcgcatagcagggactcggcagttgatgattcggttgacgatcgggtggaggtcaccactgtgacgccttctcat
 gagcccaagcacggcgagctgtga

Drosophila Gene Hit rescue sequence and BLASTX with EST and TBLASTN with
 ORF2: UDP-glucose:glycoprotein glucosyltransferase (U20554)
 20 **Human Homologue** BLASTX with UDP-GGT: hypothetical protein (AL133051)
Drosophila EST several including GH16576 (AI293351)

25 **Annotated Drosophila genome genomic segment** AE003519
Annotated Drosophila genome Complete gene candidate ugtUDP-glucose-glycoprotein
 glucosyltransferase

30 **Human homologue of Complete gene candidate** CG6850-
 IGI_M1_ctg14521_41
 D65BCE6EEC187AE3
 TRANS:SEPT20T.ctg14521.2
 2 FPC_ctg:ctg14521
 FPC_start:1284609
 FPC_end:1284696
 35 FPC_strand:+ (1.20E-215)

Putative function ugtUDP-glucose-glycoprotein glucosyltransferase

40 **Confirmation by RNAi** Only wild type profiles observed

Example 26 (Category 3)

| | | |
|----|--|---|
| 5 | Line ID | 430/5 |
| | Category | Mitotic defects in brain: metaphase arrest (overcondensation, polyploidy, metaphase with bipolar spindle) |
| | Reversion | NR |
| | Map Position | 98B5-8 |
| 10 | Rescue ID | 2C2E |
| | Rescue Sequence | GTGCGGCCCATGGATGTGCGAACGTGTACGAAGACCAAGATCGGCATCGCCA TCGGCGGCAGCACGACGGACGATAACGAAAAAGCTACAGCCGCCGCCACAGA TACAGATGCAGATGCCATGCCGCTGTTATCAGCGCGAGCGGGAGAATGATAA GGGATGGGATCGCTCAGCGCGGCAGGCAAGACGACCAAAAAGAGAGCCAAC 15 TAAATGATGTGCCTAAGACTAAGAGTTTAATGAGCATTACTGTCGCGCACTCT ATGTATTATGAATAAAATTCATACAACCTTTTGTGGTTTATTATAATAAAAGT GTGTCAGCTCTACTCGGGGGAAAGTAAGTTTACTTCTTGGCCGCTGGCTTCTTG GCGGCGACCTTCTTCTTGCGGGCGGCCAGCAACTTGGCGCGATTGGCGCAGCC TTGGTGGCCACATTGGCGAAGTGCGACTTGGCCAGCTCGACGTTCTGCTTCTT 20 GGCTTGGCCAGCACCTTGGCCACGGTGCGCTTCTCGGCGGCGAGGGCGGGCAC GACGCTTGAGTACCTCGGCATAAGGGTTCAACTTGATCAACTTGCGCACGGTT GGTAAGGGGGTT |
| 25 | Drosophila EST | several including LD45359 (AI513164) |
| | Annotated Drosophila genome genomic segment | AE003763 |
| | Annotated Drosophila genome Complete gene candidate | CG5502 RpL1 - Ribosomal protein L1 |
| 30 | Human homologue of Complete gene candidate | 1e-126 432359 dbj BAA04887 (D23660) ribosomal protein [Homo sapiens] |
| 35 | Putative function | structural protein of ribosome involved in protein biosynthesis |
| 40 | Confirmation by RNAi | Marked decrease in G1 and G2/M indicating fewer cycling cells |

Example 27 (Category 3)

| | | |
|----|---|---|
| | Line ID | 472/12 |
| | Category | Mitotic defects in brain: metaphase arrest. Meiotic defects in testis: segregation defects. Abnormal spindles |
| 5 | | (mitotic: High mitotic index, meiotic: Ab-08/24) |
| | Reversion | R? |
| | Map Position | 96C7-9 |
| | Rescue ID | 2B6E |
| 10 | Rescue Sequence 1 | |
| | | GTCTGACGTTCTCTGAGGGCAAAAGTTTCGAGTTAGTTGAAGGTGAGGGTGCT |
| | | CGATCACCGATTTGCGGTGAGACGAAAGAAAAGTATGCATTGTTGCGTTGTAA |
| | | AGAGAGCCGGCGCTCGTCTTGTTTCACATTGTCGCTGAGAACGTATGTTGTGCT |
| | | TCATCATTTTCCTTGTTGATTTCCCTTTGACGTGGCAACTTGACCATGTATGACA |
| 15 | | ACTCTTTGGTGGTGCCATCTGGAAGGCAGAAATTTGATGTCAACGGTGCTCCC |
| | | AGCCAGTCCACTCCCCAACTCACCTGCAGCTCCACTTCGATATTAACCTCGCA |
| | | ACATATTAGTGGCGTAGTTGTTCACCTGCCGCGGATCCCATTTCCGCTTTGAAAT |
| | | TTCGCACTTTCGAATATCCGTCCACATTCGATTTGAGAACATCTTCGAAACGTT |
| | | CAGCGGTGACCCAATCGGGTATTTTGCCAGCCGCCATTGTAGATAATCGGGAT |
| 20 | | AAGTATTTTGAAATCGAGCAGAAAACACATATACGTCCAGTGTGACGGTCTTG |
| | | CGTAGACTGATGAAAGCCGAGTATTAGACTCTACACATCTGTGGAGCTTTTTA |
| | | ATTCGTAGTGCGCGGCCGATTTCTCTCGATCTTCTCTCAAAAGCTCCGCTAAT |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003751 |
| 25 | Annotated <i>Drosophila</i> genome Complete gene candidate | CG10618 - novel |
| | Human homologue of Complete gene candidate | none |
| | Putative function | no homologies which indicate function |
| 30 | Confirmation by RNAi | Only wild type profiles observed |

Example 28 (Category 3)

| | | |
|----|--|---|
| | Line ID | 571/15 |
| | Category | Mitotic defects in brain: metaphase arrest |
| 5 | | (overcondensation, few anaphases, some polyploids) |
| | Reversion | NR |
| | Map Position | 93D |
| | Rescue ID | 2A8E |
| 10 | Rescue Sequence | |
| | | GGCGGCGCTACATTTGTTGTTGTCGCTGCTGCTCACAGCTCCACCACCATTTCG |
| | | ACAGTTATATTACCTCGCTCAAGTCGCCCCCTCTCCCTCTCGCCCACTCGCTGTG |
| | | TCAATCGAATTAAAACGAATGCTCTTCGGCGAATAATTGGGTTTAGATACTTT |
| | | TCCAGCAGACAAAGTTGTATTTTTTGCACCTTCTTATTGATATTAGGCAAAACGC |
| 15 | | ATCGGCCGAATCACACGCACACAAAGCACACACGCGAGCAGCGGTTTTTCAA |
| | | TCTGCAGTACACCAAACAACACACACTATTTCTTAATGCCTGTTCTTATCCCTC |
| | | TGATATTCCCAATGAATCGCTGGGCAATTGGCGATTTCGAACCGATTTTCACTT |
| | | GGCTCTTTGTTTTATTTAATTTTCACCGAAACGCTCTCACACGCAGAGACGCTT |
| | | TTGCTCGTTCGCTGATGCTTCTGCTGCAATACACACCACCTACGAAACGAGCC |
| 20 | | AAGGGAAATTGTATCTATGGGCTGTGTATCTGTTTCTACGCGGCACGCGCTGC |
| | | ACGTCCGCTCGCTTCGGGTTTTTCGAGAGAGAATATAACTTTTTTCGATACGGTA |
| | | CGGTAAACGAATTCCGCGGAATTAATTCTTGAAGACGAAAGGGCCTCGTGATA |
| | | CGCCTATTTTATAGGGTAATGCATGATAATAATGGGTTCTTAGACGTCA |
| 25 | Drosophila EST | LP07504 (AI294185), LP06548 (AI293427) |
| | Annotated Drosophila genome genomic segment | AE003734 |
| | Annotated Drosophila genome Complete gene candidate | CG15802 – novel homology |
| 30 | | to Doom, a product of the |
| | | Drosophila mod(mdg4) gene, |
| | | induces apoptosis and binds to |
| | | baculovirus inhibitor-of- |
| | | apoptosis proteins |
| 35 | Human homologue of Complete gene candidate | none |
| | Putative function | inducer of apoptosis |
| | Confirmation by RNAi | Only wild type profiles observed |

Example 29 (Category 3)

| | | |
|----|---|--|
| | Line ID | 736/15 |
| | Category | Mitotic defects in brain: prometaphase arrest (overcondensation, fewer anaphases, metaphase with bipolar |
| 5 | spindle) | |
| | Reversion | NR |
| | Map Position | 73C |
| | Rescue ID | H5E |
| 10 | Rescue Sequence | |
| | | CTAATGAGTAAGGAAAACCAATCAGCCTTGCTAATCGCTTGGCAGTATTGGCT TCTATGCAGGGGGGCGTGTCCCGCGCCCCTTGAAGCTCAAATTTTGTCAAGGG CACAGGTCGTCCCCTCCTCCTCCGCGTGGGTGGCGTTCGGCCGAACGAACCGG CGCCTACTTTGCGTCCGGCTAGCGAGGATCTCTGGGTGCCACCCACGGCTGG 15 GTGTTGCGATCTGCCCCGATTGATAATCCATGCGTGAGAAAGCTTTAGAGAATC TGCCAGATTATTACTCCCCGCATACTCAGAAAAATGTATCCTTCAGATATG TTTATGTTTATGAAGTGAAAAAAGTCCTTTGAAATACTACAAAAAGTGAGGAT CTGACCAATGATTTGATTTCTATAGAAATATACTATAAACTATAAACTAC |
| 20 | Genomic hit, Accession No. | CSC:AC014181 |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003526 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG3971 baldspot - with homology to membrane glycoprotein |
| 25 | | |
| | Human homologue of Complete gene candidate | CG3791-9e-08 4680391emb CAB41293.1 (AL034374) dJ483K16.1 (novel protein) [Homo sapiens] |
| 30 | | |
| | Putative function | membrane protein, function unknown |
| 35 | | |
| | Confirmation by RNAi | Slight reduction of G1 and G2/M peaks indicating fewer cycling cells |

Example 30 (Category 3)

| | | |
|----|-----------------------------------|--|
| | Line ID | 82/24 |
| | Category | Mitotic defects in brain: metaphase arrest |
| 5 | | (condensation, no polyploidy, no anaphases, metaphase with bipolar spindle) |
| | Reversion | NR |
| | Map Position | 100D |
| 10 | Rescue ID | 2E3E |
| | Rescue Sequence | |
| | | GGTCAAGCCCGATGGCGTCCAGCGCGGGCTCGTCGGCAAGATCATCGAGCGC |
| | | TTCGAGCAGAAGGGCTTCAAGCTGGTCGCCCTGAAGTTCACCTGGGTAAGCGG |
| | | ATAATTGAATTAGGAAGAAATCAATAGATATACATACGTGGAAACGGGGTTGC |
| 15 | | CCCACGCGGGGTTGCTATCGGACCTAACCTCAAAGGCTGGGTGCAGGCGTCAT |
| | | CGCGGAATGACATGTGTTTAGAGGTCAGAACTGCAATTAAGTATAACGAACC |
| | | GTTTTGTAACCAGGCCTCCAAGGAGCTGCTGGAGAAGCACTACGCTGATCTGT |
| | | CCGCCCCGCCCTTCTTCCCCGGACTCGTGAACCTACATGAACCTCCGGCCCCCGTG |
| | | GTGCCCATGGTGTGGGAGGGTCTGAATGTGGTCAAGACCGGTGCGCCAGATGCT |
| 20 | | CGGCGCCACCAACCCCGCCGACTCGCTGCCCCGGCACCATCCGCGGTGACTTCT |
| | | GCATTCAGGTCGGACGCAACATCATCCACGGCTCCGATGCCGTCGAGTCTGCC |
| | | GAGAAGGAGATCGCCTGTGGTTCAACGAAAAGGAGCTGGTCACCTGGACCCC |
| | | GG |
| 25 | Genomic hit, Accession No. | CSC:AC012727 |
| | Associated ORF | |
| | | Genscan ORF1 predicted sequences >16:43:49 GENSCAN_predicted_peptide_7 172_aa |
| 30 | | MKLLMLGTILAFFSVISATMAANKERTFIMVKPDGVQRGLVGKIIERFEQKGFKL |
| | | ALKFTWASKELLEKHYADLSARPFPLVNYMNSGPVPMVWEGLNVVKTGRQ |
| | | MLGATNPADSLPGTIRGDFCIQVGRNIIHGSDAVESAEKEIALWFNEKELVTWTPA |
| | | AKDWIYE |
| | | >16:43:49 GENSCAN_predicted_CDS_7 519_bp |
| 35 | | atgaagctcctgatgctcggcacaatttggcattctttctgtaatctcggcgacaatggcgggtaacaaggagaggactttcatcat |
| | | gggtcaagcccgatggcggtccagcgcgggctcgtcggcaagatcatcgagcgcttcgagcagaagggttcaagctggctgccc |
| | | tgaagttcacctgggcctccaaggagctgctggagaagcactacgctgatctgtccgcccggcccttctccccggactcgtgaa |
| | | ctacatgaactccggccccgtggtgcccattggtgtgggagggcttgaatgtggtcaagaccggtcgccagatgctcggcgccac |
| | | caaccccgccgactcgtgcccggcaccatccgcggtgacttctgattcaggtcggacgcaacatcatccacggctccgatgc |
| 40 | | cgtcgagtctgccgagaaggagatcgccctgtggttcaacgaaaaggagctggtcacctggaccccgccgccaaggactgg |
| | | atctacgaatag |
| | Drosophila Gene Hit | rescue sequence and TBLA; abnormal wing disc (awd) (X13107) |
| 45 | Human Homologue | BLASTX with awd and TBLASTN with ORF1: tumor metastasis inhibitor nm23-H2 (A49798) non-metastatic cells 2, protein (NM23B) (P22392) and nucleoside diphosphate kinase B. |

***Drosophila* EST** several including LP05977 (AI257573 similar by TBLASTX to X92956 B.taurus mRNA for nucleoside diphosphate kinase (NBR-A)

5 **Annotated *Drosophila* genome genomic segment** AE003779
Annotated *Drosophila* genome Complete gene candidate CG2210 - awd abnormal wing discs nucleoside diphosphate kinase

10 **Human homologue of Complete gene candidate** gi4505409
1A5C3F84D7AD272C
|ref|NP_002503.1| non-metastatic cells 2, protein (NM23B) expressed in [Homo sapiens] (1.90E-61)

15

Putative function human nucleoside diphosphate kinase, transcriptional regulation of c-myc expression.a candidate suppressor of tumor metastasis

20 **Confirmation by RNAi** Only wild type profiles observed

CATEGORY 4: ANAPHASE DEFECT**Example 31 (Category 4)**

| | | |
|----|---|--|
| 5 | Line ID Category | 1132/8 Mitotic defects in brain: anaphase defects (overcondensation, high polyploidy, some lagging chromosomes) |
| | Reversion | ? |
| | Map Position | 86F3-6 |
| 10 | Rescue ID | 2C3E |
| | Rescue Sequence | |
| 15 | | GGCCGGAGGTACCATTTTGGTAGGACCGTTTTTCGGGGCCAACGAAAATACCAC AAGACGGCAGCGATAATAGTGTTTTTTGCTTCAAATGTAGTATGGCTACGCAA CTCACATATGGTTAAGAAGCTTCGCTGTTTATTTGGTGGTTAACTAGCTAAATA CAATAAGAGTGGCAACGCCGTCACGTTTTCTACATGTATTTTACTTGGCGTAGT GCGCCAAGCTTATAAACCACAGTTGGGCGGTTCTTTTGAATTGTTTAATTTACA CCCCACTATGAACTTATTAGCCTTCTTTATTTATTTTATTTTATTTTATTTT AGAATACGTTTACTCAAGGTTTCGCAGCTTGTCATCAGTATTCGCAAATATCA ATAATAAAAGGCATCAATTTTCCAATCAGCAGTTGAAAAGAACTCCCCTCGAC ATTTGAACAAAATGCATTTTGGGTGATTATAATTTATTAGAATTTTATTGAC TTAAGGTAAATATAAATAAAATATTATTCAAGTACAAAGGTATATATACTCAT TAATANTATTTGGATTCAAGGAAAATATATTTCAAATGGCGGGGGTTTAATA AAACAATTTTCAAATTAAGG |
| 25 | Genomic hit, Accession No. | AC007805 |
| | <i>Drosophila</i> EST | several ESTs including LP09688 (AI295922) |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003693 |
| 30 | Annotated <i>Drosophila</i> genome Complete gene candidate | CG6929 - Lk6 kinase |
| | Human homologue of Complete gene candidate | |
| 35 | | gi4505191 DB39E49EC0BED990 ref NP_003675.1 MAP kinase interacting kinase 1 [Homo sapiens] (6.20E-113) and gi9994197 |
| 40 | | 551A82FA3D09FD58 ref NP_060042.1 G protein- coupled receptor kinase 7 [Homo sapiens] (1.70E-106) |
| | Putative function | Protein kinase associated with microtubules |

| | |
|--------------------------------------|---|
| Confirmation by RNAi cells | Complete loss of G1 and G2/M indicating fewer cycling |
|--------------------------------------|---|

| | | |
|----|-----------------------------------|--|
| | Line ID | 483/19 |
| | Category | Meiotic defects in testis: segregation defects |
| | Reversion | ? |
| | Map Position | 86F |
| 5 | Rescue ID | H2S |
| | Rescue Sequence 1 | |
| | | CTCCGGCCACACGGATGAATTCGTCGTCATTCGTCGGAATCATTTCGAACTTTG |
| | | AAAATGGATCGGTAGCTGGGAAGGAACTTAAAGCGAAATACGCAAAGAAA |
| 10 | | ACGGCTTTTGTCCGCTATTCAGCGATTTTTTTTGTGTTGTAATCAGCAGAGGAA |
| | | ATTTTAACGACCAACTCCACCGCCACACCAGCCATCTCCAGCAGCCCCGGAAA |
| | | ATAAAATAGAACTAAATTAACGCCACCATCACTACAACAACCATCTCACCAAC |
| | | AACTACAAGAGCAACAACCACAGCAACAGCACTACTGCACCAAGCCCACAAA |
| | | GAAGAGGTGAAACGCAATAATCGA=CAATACCCGAAGAAAAAAACAAAAAA |
| 15 | | ATATCGCAGATAACCGAAAAAAGCGGTGCAATAGATAAACCCCATTTTTTGCT |
| | | TGAGCTTTTTTTCGCCTGTGTGATGAGAGAAATCAGCAGCAGCCATCGATTACA |
| | | ACAACAACAGCAGCCACACCAACGACGACTCACCACCAAACGAAGAATAATA |
| | | ACCAGCGGANAGCGATAGATA |
| 20 | Genomic hit, Accession No. | CSC:AC018284 |
| | Drosophila EST | several including GH28825 (AI517767), LP04213 |

Other results same as 1132/8

Example 32 (Category 4)

| | | |
|----|---|---|
| 5 | Line ID | 1422/14 |
| | Category | Male and female sterile, small wings, meiotic defects in testis: segregation defects, elongation defect |
| | Reversion | NR |
| | Map Position | 90B4-8 |
| 10 | Rescue ID | 2F1E |
| | Rescue Sequence | |
| | | GGCCAGCTGCTCAAACATTCTGCAGCTATTTGGCCGCCAGCGAGTAGAACGAT |
| | | ATTGCCAAATATTTTATAATAGTAACCAATACGTTACCAGTATGACCGCGCCG |
| | | ATAACGATAGAAAATACCACACGGTCTAAAAGTAAATACCATTTGGGGGTATTC |
| 15 | | CCTAATCTTTTGAATTATTTACCGTTAGGTTTCGGTCGTTTTTTTTTGTTCAGCTG |
| | | TTCTTTGTATGAAACGGATTAGTAATTTTATTTGTTGTTTTTGTGCATTTTGTCA |
| | | TATTAAGCCTTGAAACATGCCTTAAATCGTTAAAATAGATTATAAGAGGGA |
| | | TGGACTGTTTGTTAAAACCAATTGGAAAATTTGTAATCGCTGGTAATAACTAT |
| | | CGAGATAAGCTTAATTATCGCTGTTTTCTTTGTATCTAGTTATAAATAATAATA |
| 20 | | ATAAACTGGTAATTAACAAAAGTAAAAAGTTACTTAAGTTATAACAAAATAT |
| | | TTAGTTATTGNATTCAATAATAAGATGGTAATAATAGATGGTAAGATAGTAAT |
| | | ATTTTAATAATTGAATTCATCACACATGCTGGTGCACGTTCCACAACCTTACAA |
| | | TCAAACGAAA |
| 25 | Annotated <i>Drosophila</i> genome genomic segment | AE003718 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG7623 - novel with homology to UDP-galactose transporter. |
| 30 | Human homologue of Complete gene candidate | 2136348 UDP-galactose transporter related isozyme 3 - human >gi 1669564 dbj BAA13527 (1e-36) |
| 35 | Putative function | sugar modification protein |
| | Confirmation by RNAi | Slightly reduced G2/M |

Example 33 (Category 4)

Line ID 1479/10
Category Mitotic defects in brain: anaphase defects
 (overcondensation, anaphase bridge, metaphase with swollen
 5 chromosomes and bipolar spindle)
Reversion NR
Map Position 69F3-7

Rescue ID 2D6E

10 **Rescue Sequence 1**

CCACGGGCAAATGTGGTCCGGAGGTCCACGACAACGTGCCGCTGACCATATC
 CCAGATTGAGCGCGCAACTCAGGATCCGGAGAACGAGAATGTGTTTCATCACA
 GACGACGTGCATCCGATTCACCTTCTGCACCTGCATCATCTACGCCTTTGTAAC
 GGCAATGGAACGCACAACGAGTCGTTTCATGAAGTTCATGATCGATGATGGCA
 15 CCGGCTCCCTGGAGGCCAGCATCACCAAAAAACCCTTCAATGGACGCGTGATC
 AGCAGCCTGTACAGTGAAGCCAGTTCGCTGGCCTCGTCCGAGGCCTACAAGA
 GCATTGCCGTGAGCATGATGCGGCTGCTGCAGGTCTCCATGGAGTACATTGAT
 CCCACGCGCATCTCGAGGGGCCACAGCCTATTCCTGCGCGGTCGTCCGAATAG
 GTTCCGCGGCAAGATGGGTCTGGACGCTTTTCAGTTCTTCATAGACAGCGGCC
 20 GATCGCGGAATATGGAAATTGGCTTCGTGGACTACCTAACCGACTGGCAACG
 AAGGCATAAAACAATGCAAAATAC

Rescue ID 2D6P

Rescue Sequence 2

25 GCCCGTGGACTTTTCACTCTGTTGATTCTTGCGTATCACGAAGTTATCCAGCTG
 GCTTTCTATGTCCTCGAACTCTGATTAAAATCCATTCTATTTGCTTAGTCTGC
 GATTTCAAAGGGGATTTCTTTATTGCAGTGCATTTTGCATTAGCGCCAAAAAA
 AAAAAAGTTGTGAGCATGGGCGTAGACTTCGTATTTTCTTACAAATAATATTA
 ATTAAAATTAATTTTGTGAGCAATTTTCACACAATTGTATTATAAGTTAAAACC
 30 AGGGTCACATTAATTTGCAGAACCGCGCAATATTTTCTTTTAAACCCCTTACA
 AATTTTCAGTTGTTTTGACTACGCCCCTGCTAATTTTACTTATTAAATTCAAA
 GTCTAAAAACATTGTCACCAGATAATACGAGTATACTATATGGACAAACGT
 AAAATCGTTAATAGAATATATATTCAACCATTATTTCAACCACCGAGAGAAA
 TTCATTTGCACAAAACGCCAGGTTGGCAGCACCATCATTGCGCACAGCAAGTG
 35 GGCAAACCTCGTTGTATCGCTTG

Genomic hit, Accession No. AC007328

Associated ORF

40 Genscan ORF1 predicted sequences >17:42:01|GENSCAN_predicted_peptide_2|1507_aa
 MKLAPTVKLNNGYEMPILGLGTYNLKKSRCEAAVCHALEMGYRHIDTAYLYRNE
 GIIGKVLAKLIGDQKLKREQVFLVTKLWDIYHEPKMVKYACDMQLKLLGVDYID
 LYLMHSPVGVDYISDEDLMPHENGQLRTNDVDYVDYRSMEQLVHLGLVRSLG
 LSNFNANQLKRLLENCQIKPANLQIECHPELVQVPLIELCKFHNTVVAYSPLGRSQ
 45 TCNPLPDYYTDSKLLALAAKYGKTPAQIILRYLSKDNEGEAAVKHAIDVGYRHID
 TAYFYQNEAEVVGKAIRDKIAEGVVKREDIFLVTKLWNIFHDPERVEGICRKQLSNF
 GLDYIDL YLMHMPVGYKYVDDNTLLPKNEDDVLQLSDVDYLDYKAMEKLVKL

GLRIEQLAGLSHLSTHSDGMQFRIRMFLTFQRGGPSHNNMQQQQQQRGGGSGTDF
YNQQRRDRRDSGRQMDNNYSNNYNNNNNNNNQRNRGGGNGMQQQQQRGGNGGSGG
GGGNGGGNNPAWNMHRGNQNSNNMMNMRNRGMGSRGPMRPNQVHLLVTHT
AIDGLLNPGFHILQGYRPQSANNQNKPRNKKIKFEGDFDFEQANNKFEELRSQLAKI
5 KVAEDGAPKPATNATAATATATNEQVGEKVEGVHTLNGETDKKDDSGNETGAG
EHEPEEDDVAVCYDKTKSFFDNISCEAAQDRSKNKKNDWRQERKLNTETFGVSS
TRRGSAHQNLNVFQAVTADATNTTTIMATAALTRDMEERQATTGTIIAWVGGGG
NFRNRSNNRNNGGGRGGNGMPNITNGNTAAALKAANNAAGHGSNATDSSAPNA
TTATTKSTSLLEQTTQVAAVSLPVLLPSIGWLFIVMDGPPDIPRSADIAILFVSFEQ
10 SVLFLKFHKRYNEFAHLLCAMMSFEDIESQLDNFVIRKNQQSEKSTGKCGPEVHD
NVPLTISQIERATQDPENENVFITDDVHPIHFCTCIYAFVTGNGTHNESFMKFMID
DGTGSLEASITKKPFNGRVISSLYSEASSLASSEAYKSIAVSMMRLLQVSMEYIDPT
RISRGHSLFLRGRPNRFRGKMGVCTNATAPSVSSINRILRNRAAERAAEFARAAS
YGYAIHPHTHPYTSFPTWPAHHPLWGAVPLATPPGGGPAGAGGALQPGGSGSSY
15 GSDGNMSSNPNSSNSNTTHSNHNTNSGSGCGDSSAGSGRLSLPALSPDSGSRDS
RSPDADANRMIDIEGEDSESQDSDQPKFRRNRRTTFSPEQLDELEKEFDKSHYPCVN
TREKLAARTALSEARVQVWFSNRRRAKWRRHQRVNLIKQRDSPSTSSSPTPLVNPV
VSPVSPVPVPVAVPESGQQKQPYPYSTSNMCNTSSSSSSNSQPCNTINPGSKMSSK
TSSVSSNQHMEEPAAAVATASPTASAPLSMGGENSAFRALPMTLPMPMTLPTASA
20 AAFALS FARQYIAKTLLGSPPRSQPPTTNQHKPEPNREFLNEACSSAASVQNSTTP
ATTADTPTAKSAMCVHCEKKGGAMEWM

>17:42:01|GENSCAN_predicted_CDS_2|4524_bp

atgaagctcgcctccgactgttaagctaaacaatggctacgagatgccattctgggcctaggaacctacaattaaagaagctcgcg
25 tgtgaggctgccgtgtgccacgccctcgaaatgggctatcgccatataagacaccgcatactgtacaggaatgaaggcattatag
gcaaggtttagctaaacttattggcgaccagaaactgaaacgcgaacagggtgttctggtcacaaagctgtgggacatataccac
gaaccaagatggtgaaatacgctgtgatatgcaattaaagctactgggcgtggactatatagatctatatctgatgcattcgccg
gtgggcgtggactacatctctgatgaagatctgatgccccacgagaatggccagctgaggaccaacgatgtggactatgtggac
acctacagaagtatggagcaactgggtgcacatctggggctgggtgcgcagcttgggattgtccaactttaatgccaatcagctgaagag
30 attactggaaaactgccaatcaagccggcaaacctacaaatagaatgtcatccggaattgggtgcaagtcccattaattgagctctg
taaatttcacaatatcaccgtgggttcctattcgccactggggcggtcccaaacctgcaatccgctgccggattactacactgattcc
aaactactggcggttggcagcgaaatacggcaagacaccagctcaaatcatcctaagatacttgtcgaaggacaacgaaggcgaa
gccgctgtgaaacatgcgattgatgtgggctatcgtcatatagatacggcctatttctacaaaacgaggccgaagtgggcaagg
cgattcgggacaagatcgcagaaggtgtgggtcaagcgagaggatataattttggtcactaagctttggaacattttccacgatccag
35 agcgcgttgagggcatttgcgcgaagcagttaagcaattttggcttggactatatcgatctgtatctgatgcataatgccagtgggcta
caaatatgtagatgacaacacctgctgccccaaaatgaggacgatgtgtcceaactgagcgatgtcgactatctggatacgtaca
aagccatggaaaagctggtaaaactgggcctgcgtatcgaacaacttgctggcctgagtcattttcaactcattcagatggcatgc
agtttcggatacggatgtttctaacattccaacgtggcggaaccagccacaacaatatgcagcagcagcagcaacgaggcgggcg
gcagtggaaacggacttctataaccagcagcgggatcgtcgggactccggacgtcaaattggacaacaactatagcaacaactaca
40 acaacaataataataatcagcgcgaatcgcgggcgggcgaacgggaatgcaacagcagcagcagaggaggaacggcgggcagc
ggcgggcgggcggtggaaacggaggtggaaacaacccggcctggaaacatgcacgcgggaaccagaactcgaacaacatgatg
aacatgcgcaaccgcggcatgggatcccgcgggcccatgcgaccaatcaggtacacctgctgggtgactcacactgctatagat
ggtttattaaacctgggctttcacattttgcagggctatcgtccgcagtcggccaataatcagaacaagccgcggaacaagatcaa
gttcgagggcgacttcgatttcgagcaggcaacaacaagttcgaggaactgcgctcccaactggccaagctcaagggtggccga
45 ggatgggtgcaccaagccagccaccaatgcaacggccgccactgcaactgcaaccaatgagcaggtgggtgagaaggttgaa
ggcggttcacacactgaatggcgagaccgacaagaaggatgattctggcaacgagaccggcgctggagagcacgagcctgagg
aggatgatgttgctgtgtgtctacgacaagaccaaactcgttcttcgacaacatctcgtgcgagggtgccaggatcgagcaagaa
caagaagaacgattggcgccaggagcgcaagtgaacacggagaccttcggagtgctctccacacgacgtggcagtggtggctc

atcaactgaatgtattccaagcagttaccgcgacgcaaccaataactacaacaataatggcaacggcggcattaactcgggatatg
 gaggagcggcaggctacaacaggaacaattatcgcatgggtggcgggcggaacttcgaaacaggagcaacaatcgc
 aacaacggcggcggtcgtggcggaacggaatgccaacatcaccaatggcaacacggctgctgcgctgaaggcggccaac
 aatgctgctggccacggatccaatgccacggactccagtgcacaaatgccacaaccgcgacgacaaaagtcgacgtccctcttg
 5 ccagagcagacgcaacaggtggcgggcagtttcgttggccgtgttggtaccatcgattgggtggcttttatcggtatggatggaccac
 cagacattccaagatcggcagatattgcgattctcttcgttagtttgaacaaagtgtacttttcttaatttcacaagcgatacaacg
 agtttgcccacttgctgtgcgcaatgatgagtttcgaggacatagaaagccagctggataacttcgtgatacgcaagaatcaacag
 agtgaagtcacgggcaaatgtgttcacacagacgacgtgcacccgattccttgcacctgcatctacgccttctgaactgg
 ggatccggagaacgagaatgtgttcacacagacgacgtgcacccgattccttgcacctgcatctacgccttctgaactgg
 10 caatggaacgcacaacgagtcgttcacgaagttcatgatgatggcaccggctccctggaggccagcatcaccaaaaaacc
 ctcaatggacgcgtgatcagcagcctgtacagtgaagccagttcgtggcctcgtccgaggcctacaagagcattgccgtgagc
 atgatcgggctgctgcaggtctccatggagtacattgatccacgcgcacatcgcagggggccacagcctattcctgcgcggctcgc
 cgaataggttcgcgggcaagatgggtgtctgcaccaatgccactgctccttcgggtgagcagcatcaatcgcatattgcgtaatcga
 gcggcggaaggcagctgcggaatttgcctggcgggcaggttacggctatgccatccacacacatccgcacatccgtacacc
 15 agtttcccacttgccggcgcatcctcgtgtggggagccgtgcccctggccacgccacctgggtggcgccctgctggagcc
 ggtggtgactgcagccggcgggcagtgccagcagctatggcagtgatggcaacatgagctcaaatcccaatagcagcaaca
 gcaacaccacccacagcaatggccacaatacaacagcggcagtggtatgcggggatagtagtgccggaagtggacgcctctc
 cctgccggcactttcgcggattccggaagtagggacagccgctccccagacgcagatgccaatcggtgatagatatcgaagg
 cgaggacagcagtcgcaggacagtgaaccagccgaagtccggcgcaatcgcaccaccttcagtccggagcagctggatgag
 20 ctggagaaggagtgcgacaagtcgactatccctgcgtgaatacccgcgagaaactggccgcccggacggcactgagcaggg
 ccagggtgcaggtttggtttccaacagacgagcgaatggcgggcgccaccagcgggtcaactgatcaagcagcgcgactcg
 ccctcgacatcgagctcaccacgcggttggtcaatccggtggtcagtcgggtcagccaatccagttccagttccagttgcagtt
 ccagaatctggccaacagaagcagccatatccgtacagcaccagcaacatgtgcaacaccagcagcagcagcagcaacagtc
 aaccgtgcaacaccatcaatcccgcgagcaaaatgagcagcaaaaccagcagcgtcagcagcaaccagcacatggaagagc
 25 cagcagcggcggtggccactgcctcaccacagcatcagctccattatcaatgggcggtgagaacagtgcatcttcgcgctctgcc
 catgaccttgccgatgcccacacgttgcccacggcagtcggcgggcgcccttcgcgctcagcttcgcccggcagtcacatagccaa
 gacgcttctcggttctccagatccagatccagccaccaaccaccaaccagcataagcccagccaaatcgcgagttcctcaat
 gaagcctgcagctccgcagcatctgtccagaattcgacaacgcgggcaacaaccgcagatactcctacagccaaatcagcaatg
 30 tgcgtgactgcgagaaaaaggaggggccatggagtggatgtga

***Drosophila* Gene Hit** BLASTN with rescue sequence 2: Histone acetyltransferase GCN5
 (AF029776) very small match at end, TBLASTN with ORF1:
 middle domain histone acetyltransferase GCN5 (AF029776).
 Genomic matches histone acetyltransferase

Annotated *Drosophila* genome genomic segment AE003541
Annotated *Drosophila* genome Complete gene candidate CG4107 -Pcaf /GCN5 histone
 acetyl transferase
 transcriptional activator
 protein
Human homologue of Complete gene candidate gi6382076
 72F516F8BD10CD0C
 [ref|NP_003875.2| p300/CBP-
 associated factor [Homo
 sapiens] (1.20E-197)

Putative function Transcriptional activator

Confirmation in RNAi Only wild type profiles observed

Example 34 (Category 4)

Line ID 184/5
Category Mitotic defects in brain: Anaphase defects.
5 (overcondensation, aneuploidy, some lagging chromosomes and breaks)
Reversion R
Map Position 71B

10 **Rescue ID** C4E
Rescue Sequence
CTCGAGCAGATGTGGGACGAGCTGAGCGGAGCGCACAAACTGCCAAGTAAGT
GGAGCATGTGGATGAAAGGAGTTCCCAGAACAGTGTTGCCAACCAAAAAAAAAA
AAAAAAGTTAAAAAGTTAATTTTAATAGTGTAATAAATATGAATTAAATTAA
15 ATTTTTATGTAAACAGTATTAGCTTTACATGAGATTACCAAATTGTGAGTGTCT
GTGTTTGTGTCTTTTAAAAACTTTAAAAGCACATAAAGAAATATATTTTAAA
TTTAATTAAAAAGTTCGTAAAAAGTAAAAGGTAGCTAAATTAAAAAGTTTCCT
ATTCAAATCAGATTTGGCGAACAAAGAGCCAAGTTGGCAACACTGACAATGA
CTCCAAGCGCGAACAAAGCGATTTCTATCGTTATCCCACTCTCTCTCCAGAG
20 ATCGTTCTCAAGGCCAAATGGAAGGGACTTCGAGACAATTTCCGTGTGGAGTC
AAAAGGATCCGGCGGCCGAATAACGG

Genomic hit, Accession No. CSC:AC019852

25 **Associated ORF**
Genscan ORF1 predicted sequences >22:43:26|GENSCAN_predicted_peptide_2|1003_aa
MAPKKSTIVLNVEQFIHDIERP AIWNRNFHCNKAFLEQMWDLSGAHKLPKIVL
KAKWKGLRDNFRVEYKRIPRADNGDFMVDPATFESKWLHYYALLFLTDHMRHR
LPKNEQDQSFYFSQQSEDCEKT VVEPDLTNGLIRRLQDSDEYDEEEMEADGEAS
30 EATMEETMPTPPAAHQMNQVSTTPLATGALRAQEEAHQHALLKAGLLRAQLMEL
EKEAEDLSRKPPPPQQMTSPVAPSLQVLVEPPAAHCSPPPMVTTTSAQVQQPGSA
AVLAPATTTSSASSVSSNGAPMGGKRSVSPPLYNKAHHPLATLAAHLAAKDRN
EDFGPTSAVGGNGDHLSFTQHSYANGLIPALKLKRPRLSNDFNGSSTMDTPLVP
EDDDYHYLLSLHPYMKQLTAAQKLRIKTIKLIKFELYKEDLEESNLDREVYVL
35 DDGAEVDLDLGNYERFLDVTLHRDNNITGKIYKLVIEKERTGEYLGKTVQVVP
ITDAIQEWVERVAQTPVQGSSKPQVCIVELGGTIGDIEGMPFVEAFRQFQFRVKRE
NFCLAHVSLVPLPKATGEPKTKPTQSSVRELRGCGLSPLIVCRSEKPIGLEVKEKI
SNFCHVGPDQVICIHDLSIYHVPLLMEQNGVIEYLNERLQLNIDMSKRTKCLQQ
WRDLARRTETVRREVCIAVVGKYTKFTDSYASVVKALQHAALAVNRKLELVFIE
40 SCLLEEETLHSEPSKYHKEWQKLCDSHGILVPGGFGSRGMEGKIRACQWARENQ
KPLLGICLGLQAAVIEFARNKLGLKDANTTEIDPNTANALVIDMPEHHTGQLGGT
MRLGKRITVFS DGPSVIRQLYGNPKSVQERHRHRYEVNPKYVHLLLEEQGMRFVG
TDVDKTRMEIIELSGHPYFVATQYHPEYLSRPLKPSPPFLGLILASVDRLNQYIQRG
CRLSPRQLSDASSDEEDSVVGLAGATKSLSSLKIPITPTNGISKSCNGSISTSDSEGA
45 CGGVDPTNGHK

>22:43:26|GENSCAN_predicted_CDS_2|3012_bp

atggcgccaaaaagtccaccattgtgctcaatgtggagcagttattcacgacatcgaggagcgcccgccatctggaaccgca
 atttccactgcaacaaggccttcctcgagcagatgtgggacgagctgagcggagcgcacaaactgccaaagatcgtgctcaagg
 ccaaatggaagggacttcgagacaatttccgtgtggagtacaaaaggataccgcgggaggataacgggtgatttatggatcc
 5 ggccacctttgagtcgaagtggctgcactactatgcattgtgttttaactgatcacatgcgtcatcgtttgccaaagaacgaacagg
 atcagtcattttacttcagccagcaaagcgaggactgtgaaaagacagtgggtggagccggatttaacaaacgggtctaatacgtcgt
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 gatgccacgcccacggctgcgcataaatgaatcaagttagcaccacaccactggccaccggagctttgcgagcccaagaag
 aggcacatcagcacgctttaattaaggcaggattactccgcgctcagttgatggagctggaaaaggaggcggaggacttgagca
 10 gaaagccacctccgccacagcaaatacatctccagtggcaccctcactacaagtgtatgtggaaccaccagccgcacactgtt
 ctccaccgccaatggtgaccaccacatccgcacaagtacaacaaccgggctcagcagctgttctggcgccggcaacgaccaca
 tccgctcatctgtatcctcgaatggagcgccaatgggaggcaagagatctgtgtcgcaccgcctctatacaacaagcacacc
 atccgctggccactctggcagcagcacatcttgcggccaaagaccgaaatgaggatttcggaccacactctgtgtaggaggaa
 acggagatcacctgagcttcaactcctacgccaatggactgatacccgcccttaagctgaagcgcccgctctctccga
 15 ggatagcaattttaatggttctcgaatggacactccgctcgtaccagaggacgatgactaccactacttgcagcctacatcc
 gtacatgaagcagctgaccgcagcccagaagctgcgcatacgcaccaagatacaaaaagctcatcttcaaggaaactctacaaga
 agatcttgaggagtcgaacctagatcgcgaggtttacgttttgacgatggcgccgaggtggatctggatctgggaaactatgaac
 ggttttgatgttacctgcacgggacaacaacataaccaccggaaaaatttacaagttggtcattgagaaggagcgcactggc
 gactacttgggcaaaacgggtcaagttgtcccacacatcactgatgccattcaggaatgggtggagcgcgtggccagacaccc
 20 gttcagggatcttcaaagccacaggtgtgcacgtggaattgggaggaaacgattgggtgacatcgaaggcatgccttctgtagagg
 ccttccgtcagttcagttccgcgtaaagagagagaacttctgtttggcccatgtgtcgtggttccgttgccaaaggctaccggag
 aaccaagaccaagcccacacaaagttcggtcagagaactgagaggatgtggcctgagtcctgattgtctgccgatcggg
 gaaaccattggactggaggtcaaggagaagatcagcaactttgtcatgtggggccggatcaggtgatatgcacccagattga
 actccatttatcatgttccgctgctgatggagcagaatggtgttattgaatacctaaatgagcgcctacagcttaatatcagatgagc
 25 aagaggaccaaatgcttgagcaatggcgagatttggcgcgctgaacggagaccgttcgccgtgaagtttgcacgccgtcgtg
 ggaaagtacaccaagttcacggattcgtacgcctccgtagttaaagccctgcaacatgccgcctggcagtgaaatcgcaactgg
 aactggtctttatcgagtcgtgcctgctggaggaggaaactttgcattctgagccgagcaagtagccacaaggagtggcagaagct
 atgcgatagccatggcatcctagtcctccgggtggattcgggtccgtggaatggaggggcaagattcgtgcatgccaatggcgcgga
 gagaatcaaaagccattgcttggcatctgcttgggtctgcaagcggcggtcattgaattcgacgaaataaacttgggtctcaaggat
 30 gcaaacaccacagaaatcgatccgaacacagctaataccttggatcgcgatatgccagagcatcacacgggtcaattggcgggc
 actatgcgcttgggcaagcgaataactgtttctctgatggtcctagtgatcattcgccagttgtatggcaatccgaaaagcgtgcagg
 agcgtcatcggcatcgttacgaggttaatcccaatacgtgcacatcgtggaagagcaaggcatgcgatttggggcaccgacgt
 cgacaaaactaggatggaaatcattgagctcagcgggtacccctactttgttgccaccaataatcatccagagtacttgcgcggcc
 tetgaagccgtgcctccttctcctggcctgatcctggcctcagtgatcgattgaaccaatatattcagcgcggttgccgcctgtcg
 35 ccccgccagctatccgacgcacatcctcgatgaggaggacagtgttgggttggccggagcaacaaaatcgctgagctccttg
 aaaattcccattaccccacaaatggaatatcaaaaagttgcaatggtagcataagcacttccgacagcgaaggtgcctgcggag
 gcgttgatcctaccaatggccataagtaa

40 **Human Homologue** TBLASTN with ORF1: CTP synthase (CTPS) (NM_001905.1)
Drosophila EST LD27370 (AA941993)

Annotated Drosophila genome genomic segment AE003532
Annotated Drosophila genome Complete gene candidate CG6854 - novel protein,
 possible CTP synthase?

45 **Human homologue of Complete gene candidate** gi4503133
 C33BD849A0044697
 |ref|NP_001896.1| CTP

synthase; cytidine 5-prime
triphosphate synthetase
[Homo sapiens] (8.40E-217)

5

Putative function Enzyme important in the biosynthesis of phospholipids and
nucleic acids, and plays a key role in cell growth, development,
and
tumorigenesis. The region of the human gene is the location of
breakpoints involved in several tumor types

10

Confirmation by RNAi Loss of G1 and G2/M peaks indicating fewer cycling cells

Example 35 (Category 4)

5 **Line ID** 225/27
 Category Meiotic defects in testis: segregation defects
 Reversion NR
 Map Position 90D

10 **Rescue ID** 2D2P
 Rescue Sequence 1

Rescue ID 2D2E

15 **Rescue Sequence 2**
 GCCTGAACTTAAAACGCTGCCTTCGGCTCTCGCTCGGCACTCGCTCGGCTGCG
 ACGTCGACTGCGACGCTGGCAGCGACAACAACGATTGGCCTCTCTCATTCACT
 TACCTCCTCTCTCTCTCTCGCACTCTCTCTTAGCGGTGAGAGAGTGTTTTCTC
 ACATTTGTTTTGCTTTTGCGGTTTCGCCAATGGCCCCCAGAAAAGAGCG
 20 CGCAAGAGCTAGCTCCACAGTGGATCCTAAGAGAACGGTCCCTGTGGACTCC
 ATCTAGCTAAGAGAAACGCACTTAGTTAGTTTCTATTTTTGGTTGTTTAAGTAC
 TGCTAGCTGCCTGCCAGTTGAGTGTCCGTCCAAAAACGGTGGTGGAAATGGGG
 GTGACCACTTCAAACATGAAAGCGAAATGTCCTGAGACCCTACAAAACTAG
 AAATACGCGGGTGCCTGAGAGAAATTTTTTATTTCAGTAAATTGGCAGAGG
 25 CTACATTTTGAATGTTTACAATGAAAATTGCTGGGGGAAGCTAGTGAACAACCA
 TTTCGCCATAATTACACTATCTAAGCTTTTATTTTTAGCCACATGATATATGC
 ATGCA

30 **Genomic hit, Accession No.** AC008361

Associated ORF
 Genscan ORF1 predicted sequences >20:36:39|GENSCAN_predicted_peptide_2|515_aa
 MSSTIRLQTSSCQCKLYKYERHPNKP NLQPTPIPNYPCEILHIDIFALEKRLYLSCI
 35 DKFSKFAKLFHLQSKASVHLRETLVEALHYFTAPKVLVSDNERGLLCPTVLNYLR
 SLDIDLYYAPTQKSEVNGQVERFHSTFLEIYRCLKDELPTFKPV ELVHIAVD RYNT
 SVHSVTNRKP ADVFFDRSSRVNYQGLTDFRRQTLEDIKGLIEYKQIRGNMARNKN
 RDEPKSYGPGDEVFVANKQIKTKEKARFRCEKVQEDNKKNRNGKAAGGKGKTR
 RVARGAQIYQNWAICRNLF LFLSLACCRVCKVCDIVVEFRKGTNAV VNVQIREAI
 40 SHVFHKEDIVIDVQESKEWCIWTDDQVQSPLPELENLWHELWIGPSHAYLIDQIVD
 LFENLLEKYNVQVVDVVRFNFLHRA LVVVIISGIIIIIMIIGVSGGQRTNAFSHRS
 QRSAIGGDPQKDS AVQQVQARSSDAFCQIPHRS PRFPGRSQLIPKPNREILRNASA
 TKNLLFRIRSQ

45 >20:36:39|GENSCAN_predicted_CDS_2|1548_bp
 atgtccagtacgatccgtctgcaaacttctcatgtcagtggtgcaaactctacaagtacgagagacaccctaacaaccaaaccta

caacctacgccaattcctaactacccatgtgaaatacttcacatcgacatttttgcgctcgaaaaaagggtatacctaagttgtattgac
 aaatttagcaagtttgccaaacttttccatctgcagtcaaaagcatctgtgcatttgcgagaaactttggaggaggccctacattacttc
 accgcccctaaggtcttggtttcggataacgagcgagggttggtatgccccacagtgtcaactatcttcggtctctagatcgcgatc
 gtattatgctccaaccagaagagcgaagtaaatggtcaagtcgagagattccactctacgttcctagaaatttatcgttgccttaaa
 5 gatgagctccctaccttcaaaccggtgagctggtacacatagcagtgaccgctacaacacttccgttactcggtaacgaatcg
 aaaaccagcagacgttttttcgaccgctcgtcaagggttaaactatcagggtctgacagatttccggcggcagactttagaggacat
 caagggttaattgagtataagcaaattagaggtaatatggctcggataaaaaataggagcagagccaaagtcttatgggcccggga
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 15 aagaacaaatgccttttcacaccaccgatctcagcgatcagcgatcggcggcgaccctcaacaaaaagattcagcgggtgaaca
 ggtgcaggcacgatcttcggatgccttttgccagataccccaccgatctcccagggtcccaggggcgagccaacttattccgaagc
 caaatcgagaaattcttcgaaacgcgagtgccacaaaaatttattgtttcgaattcgcagccagtga

20 ***Drosophila* Gene Hit** BLASTN with rescue sequence: couch potato (Z14974).
Human Homologue BLASTX with couch potato: RBP-MS/type 2 (RNA binding motif family)(D84108)

25 **Annotated *Drosophila* genome genomic segment** AE003720
Annotated *Drosophila* genome Complete gene candidate CG18434 -couch potato RNA binding protein
 30 **Human homologue of Complete gene candidate** 2224621 dbj|BAA20798| (AB002338) KIAA0340 [Homo sapiens] (2e-19) and Ensembl predicted peptide Gene:ENSG00000070877 Clone:AC009710
 35 Contig:AC009710.00004 (predicted unknown protein)

Putative function Possible RNA binding protein

Example 36 (Category 4)

| | | |
|----|-----------------------------------|--|
| | Line ID | 238/37 |
| | Category | Meiotic defects in testis: segregation defects, multi-stage defects |
| 5 | | (P1-02/17) |
| | Reversion | ? |
| | Map Position | 70D |
| | Rescue ID | I7E |
| 10 | Rescue Sequence | |
| | | GTTCAAACGCACTTTTAAGGTGGCCTATCGGCCCATCAGGAGCAACTTTGTTC |
| | | TGCTGCCGGATCAGTACTACGGCGTCGTTTCGACCTATGTAAGTGTCTAAAGG |
| | | TCTTCGCTCCATTCAGATTAGATACGCCAAAGATTAATCCGGTCAACCATTCT |
| | | GATTAGGACACGGGCTGCCTGAGCTTGCAGTACAATGGTCGGACGCACTACG |
| 15 | | CCTCCTGGGCGCCACAAAAGGGCGGGCGGCATTAAAGACACCGAGATTGG |
| | | GATCAATGCCAGAGCGGCCAAGGAGATCGGTAAGCCATTACTTAACGGCCGG |
| | | ATGTGCATCGGTTGCCAATGTGCCGTAATATTGGACTCCGGCCATCTGCCCCG |
| | | TACCTCGTACGCTAGCAGCACCCACTTACCCTTTCTTGCCGTAGGTCTGCACGA |
| | | GAATGATCTGGTCAAGTGTGCGCTCATCGCTGACGTTCTCAACCTGCGCAGCG |
| 20 | | TCCACGTTACCCCCGTCTCGTCCAAGGACTGGGAGATCATAGTGAGTGACGGT |
| | | TTCGCCTGCTTGGCGGGCGTGG |
| | Genomic hit, Accession No. | CSC:AC017664 |
| 25 | Associated ORF | |
| | | Genscan ORF1 predicted sequences >15:26:30 GENSCAN_predicted_peptide_1 1819_aa |
| | | EMVQAKDPPSHYLSKLRTYLDPKASRSHRLYLFYFLCQKRKMVGESTSTQVLRD |
| | | LEISLRTNHIEWVKEFLDDTNQGLDALVDYLSFRLQMMRHEQRLQGVLCASEERL |
| | | NLTNGGDGGEIVMGNSSSVSPGGGGGLLSHGNSTGHGLANGTLDNRQQHTMSYG |
| 30 | | FLRPTIADALDSPSLKRRSRHIAKLNMGAAATDDIHVSIMCLRAIMNNKYGFNMVIQ |
| | | HREAINCIALSLIHKSLRTKALVLELLAAICLVKGGHEIILGSFDNFKDVCQEKRRF |
| | | QTLMEYFMNF EAFNIDFMVACMQFMNIVVHVSVEDMNYRVHLQYEFTALGLDKY |
| | | LERIRLTESEELKVQISAYLDNVFDVAALMEDSETKTSALERVQELEDQLEREIDR |
| | | NSEFLYKYAELESESLTKTEREQLAMIRQKLEEELTVMQRMQLQHNEQELKKRDT |
| 35 | | LLHTKNMELQTLRSLSLPRSASSGDGSLANGGLMAGSTSGAASLTLP PPPPPMPASP |
| | | TASSAAPPPPPPPAPPAPPPPPGFSPSGSLASTAPSPPHAPPMLSSFQPPPPPPVA |
| | | GFMPAPDGMATIKRKVPTKYKLPTLNWIALKPNQVRGTIFNELDDEKIFKQIDFNE |
| | | FEERFKIGIGGALRNGSNGTEVDGSLQSSKRFKRPD NVSLLEHTRLRNIAISRRLG |
| | | MPIDDVIAAIIHSLDLKKLSLENVELLQKMVPTDAEVKSYKEYIIERKDQQLLTEED |
| 40 | | KFMLQLSRVERISSKLAIMNYMGNFVDSVHLISPQVQSIAGASTSLKQSRKFKAVL |
| | | EIVLAFGNYLNSNKRGPAYGFKLQSLDTLIDTKSTDKRSSLHYIVATIRAKFPELL |
| | | NFESEL YGTDKAASVALENVVADVQELEKGMDLVRKEAELRVKGAQTHILRDFL |
| | | NNSDKLKKIKSDLRHAQEA FKECV EYFGDSSRNADAAFFALIVRFTRAFKQHD |
| | | QENEQRLRLEKAAALAASKKENDQVLMRNKVNQKKQQA VINELKSKAHSVRE |
| 45 | | KKLLQQDEVYNGALEDILLGLKSEPYRRADAVRRSQRRRIDNNRLSRTLEEMDCL |
| | | HENDLVKCALIADVNLRSVHVTPVSSKDWEIIEI LSTEKISGSVLEQTRIVNSTQILI |

VWINKSMQVALTVDRLKPHMNYGRIDHNTELVVAPNLYKGLTNGTSNGVIEENT
 KLSRSKTTAQVKDELTEKLTPALTHSSTVSNVKNTIQRNKRQDHMERLKKDLRRES
 SRSFEFRVIRGLWREQAQESDVFNKGHLPEFFDLDFYCMHTAADKDYYVRVR
 TVEDDIEDDLPETIHPSIELNANLMKLLGIKELERVVLRPKTTVVNFVEKIELFANK
 5 KTHYKIMENAFKRFVIERTQHKPMLFNQEEVVRLEDDLLVTVGILPEHFRYCVVD
 AQFLKESKIYAADLVRPVGEIIEETPPTSPLSVQDLIQLPEYDKIVDQVVQELRMN
 LCLSADNSVMRQCNVLLAGASGTGKTVLVERILDQLSRKPDYCHFEFFHGSRSKG
 RKTESIQKDLRNIFTSCLOHAPAVVLENLDVLAHAAGEQSSQDGEYYNRMADTV
 YQLIVQYTTNNAIAVIATVNELQTLNKRLLSSPRGRHVFTVARLPNLERADREILR
 10 ELCSHINVAKDLDLVKFSNLTEGYRKCDLVQFVERAIFYAYRISKTOPLLNTNDQLI
 ESLEHTNSYCLQGIQSNQRTGNDADANEMRVEELPGLESVVGVLVLMWPSRY
 PTIFNASPLRNQAGVLLYGPPGTGKTYLVSQALATSWNLRIISVKGPELLAKYIGQSE
 ENVRNLFNRARSARPCVLFDEFDSLAPKRGHGSTGVTDREV

15 >15:26:30|GENSCAN_predicted_CDS_1|5457_bp|
 gaaatggtgcaggcaaaggatccgccctcacattacttgagtaaactgcgcacatatctggacccaaaggcatcaaggagtcac
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 20 ggcgatggcggtgagatagtgatgggaaacagtagttctgttagtcctgggtggaggtgggtggttactatcacatggaaacagtac
 gggacatggctggccaatggcacactgactcgaggcagcagcacacaatgtcctatggattcctacgacctaccattgccgat
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 agtcttatccacaacatcgctgaggacgaaagccctggctcctggagctgctggcagccatctgtctggttaaaggaggacacgaa
 25 atcattttgggttcgttcgataatttaaggatgtgtgccaggagaagcgcgcttccaaacgctcatggagtactttatgaacttcga
 ggctttaaataagatttatggttgctgcagtcagttcatgaacatcggttgcactcgggtggaggacatgaactacagggtgcac
 ttacagtacgagtttacagccctgggcttgataagtatctggagcgaattcgattgacagaatcggaggaaactgaaggtgcagat
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 gcttgaggatcaactgagcagagaaatagatcgttaactcagagttcctctataagtatgcggaattagagtcgagagctaacgt
 30 gaaaacggaacgcgagcagctggctatgattcggcagaagctggaggaggaaacttacagtgatgcagcgaatgttgacgacaca
 acgagcaggagctgaagaaacgggacacactgctgcacacaaagaacatggagctgcagacgcttccggttcctgccacga
 tccgcctccagcggcgatggttcttggcgaatggtggcctcatggctggttccacatcgggggcagcctcttaacattgccacc
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 accaccgccgggttcagtcgctgggcagtcgagcggcagcctagcctcgacagcggccatcgccgccacatgccccgcc
 35 atgctaagctccttccaaccgccaccgctccagtgccggctttatgcccgtcccgatggcgccatgacctcaaacgcaagg
 tgcccactaaatacaagttgccacctgaactggatagcactaaagcctaatacaggtacgtggtacaataattcaacgagctggatg
 acgaaaagatcttcaagcaaatcgacttcaatgagtttgaggagcgttcaagatcgggattggcggtgctttgcgcaatggttagc
 aatggaaccgaggtcgatgggtcgctgcagtcagcaaacgcttcaagaggcccgacaatgtctcgctgctggagcacacgag
 gtttaagaaacattgcaatctcccgtcgcaagctgggtatgccattgatgtcatcgccgccattcatagtctggacctgaagaa
 40 actttccctggagaacgtcgagctgctgcaaaaaatggtgcccacggatgccgaggtcaaactcacaaggaatatatcatcgag
 cgcaaggaccaacagctactaccgaagaagacaagtttatgctgcagttgctgcgtgtggagcgtatctcgtccaagctagcca
 ttatgaactatatgggcaattttgctgacagcgttcattatgctccgcaagtgaatcgatagcaggagcgtcgacttccctaaaa
 caatctcgaataatcaaggcgggtttggaaattgtcctggcttcggcaactatctcaacagcaacaaacggggaccagcctatgg
 cttaagctgcaatcgctggacacgctgatcgatacaaaatccacagacaagcgtcgtcactgcttactatattgtggccaccat
 45 acggggccaaatttcggagctgctgaacttcgagagcagctgtatggaacagacaaggctgcacgggtggcactagagaatgt
 ggtggccgatgttcaggagcttgaaaagggcagtgatctggtgcgcaaggaggccgagctgcgagtgaaagggtgccagacg
 catatcctgcgtgacttctgaacaacagcagggacaagctgaagaagatcaagagcgtatcgggcatgcacaggaagcgttc
 aaggagtgcttgagtactttggcgactcctcgcggaatgcagatcgggctgcttcttgcgttgatcgcttcacgagagcgc

ttaagcaacacgatcaggagaacgagcagcgcttctgcctggaaaaggccgctgcgctggccgctccaagaaagagaacga
 tcaggtgcttatgcgcaacaaggtaaccagaagaagcaacaggaagctgtcataaacgagctgaagagcaaggcgactcgg
 tgcgcgagaaaaagctgctgcagcaggacgaggtgtacaacggagccctggaggacatcctgctcggcctgaagagcgagcc
 gtacaggcgggcggtatgctgtgcggcggtcgcagcgccggaggatcgacaataatcgtttatcgcgcaccctggaggaaatgg
 5 attgtctgcacgagaatgatctggtcaagtgtgcgctcatcgctgacgttctcaacctgcgcagcgctccacgttaccctcgtctcgt
 ccaaggactgggagatcatagaacttagcactgaaaagatatcgggcagtgtgctggaacaaactcgcatagtgaattcaacgca
 gatccttattgtttggattaataagtcgatgcaagttgcgctgacagtggatcgctgaagccgcacatgaactacgggagaataga
 tcacaatacggaaactcgtggtggcgcccaatctgtacaagggtctgaccaatggaacttcaaatggtgttatagaggaaaacacaa
 aactctccagaagtaaaaccactgccaggtcaaggatgagctgactgaaaagttaacaccgttgacccattcctccacggtgtcc
 10 aatgtgaaaaatactattcagcgtaacaagcgtcaggatcacatggagcgtcttaaaaaggacttgccgcgcgaaagctcgcgta
 gcttcgaatttcgtgtcattcgaggtctatggcgggagcaggcccaggagtcggatgtgtttgtgaacggaaagcatctgcctgag
 ttctttgatctagatctattctattgcatgcacaccgcagccgacaaggattactatgtgagagtgcgcacagtagaagacgatattg
 aggacgatctaccagaaaccattcatccatcgatcgaactaaatgccaatcttatgaagttgctgggtattaaggaattggaacgag
 tggttctaagacctaataaactaccgtagttaactttgtagaaaaaattgagctatttgccaacaagaagacgcactacaaaatcatgga
 15 gaacgcatttaagcgatttgtgatagagagaactcagcacaagccgatgctcttcaaccaggaggaggtggtacggctggagga
 cgatttactggttactgttggaattttaccagaacactttcgttattgcgtggtggacgcgcagtttctgaaggagtccaagatctacg
 cagcagatctggtgcgtccggttggcgagattattaaggaggagacgcctccgacatcgccactaagtgttcaggatctcatcca
 gttaccggagtacgataagattgtggatcaggtagttcaggaattgcgaatgaatctatgcctcagtgcgtgacaattccgtcatgcgt
 cagtgcattgtcctactcgtggtgcctcgggaacgggtaaaacagttctgtggagcgcattttggaccagctgtcacgcaagcc
 20 ggattattgtcacttcgagttcttccacggatcgcgaagcaaaggccgcaagacggagtcctccaaaaagatcttcgcaacatttt
 taccagctgcctgcagcatgccccgcctattgttgtctagaaaacttggtatgtactggcccacgctgctggagagcagtcacgtc
 aggtatggagagtactacaatcgcatggcggatactgtgtatcagttgattgttcagtataccaccaacaacgctattgcagtaatcg
 ccaccgtcaacgagttgcagaccctcaataagcgattgagctcaccaaggggaagacatgtcttccagactgttgctcgtctgccc
 aatttggaacgagcagatcgagagataattcttcgagagctgtgcagccataatgtggccaaggacctggatctgttaagttct
 25 ccaacctcacggagggtaccggaaatgtgatcttgttcagttcgtggagcgtgcaatattttatgcttatcgcataagcaagaccc
 agcctcttctgaccaatgatcagcttattgagtcctggagcacacaaactcgtactgcctgcagggcattcagagcaatcaaaga
 actggcaatgatgccgatgccaatgaaatgcgcgtcgaggagttgcctggcctggagtcagttgtgggagttctggaggaggtcc
 ttatgtggccatcaaggtatccaaccatttttaacgcctctccactgcgcaaccaggccggagtacttctatatgggccaccaggaa
 caggtaaaacctatctggtctctcagttggccacatcgtggaacctgcgcacatttccgtaagggtcctgagttgctcgccaaata
 30 tattggtcaaagcgaggaaaatgttcgaaacctgttcaatcgagctcgcagtgcccgaccatgtgtgcttttctcgacgagtttgac
 agcttggcgccgaaacgtggtcacgattccacgggggtcaccgatcgagtg

Drosophila Gene Hit recue sequence and TBLastn with ORF1: mRNA for l(3)70Da (AJ243811)

35 **Human Homologue** BLASTX with l(3)70Da: peroxisome biogenesis factor 1 (AF026086)

Drosophila EST LD43687 (AI512050)

40 **Annotated Drosophila genome genomic segment** AE003536

Annotated Drosophila genome Complete gene candidate CG6760 mRNA for l(3)70Da
 - novel protein with
 homology to endoplasmic
 reticulum ATPases

45

Human homologue of Complete gene candidate 4505725

ref|NP_000457.1|pPEX1|
peroxisome biogenesis factor
1 >gi|2655141 (AF026086)
(8e-80)

5

Putative function Putative member of the AAA protein family (ATPases associated
with diverse cellular activities) including homologies to
transitional endoplasmic reticulum atpases, and an E.coli
membrane-bound AAA-type metalloprotease which degrades
degrades sigma32, an alternative sigma factor for heat shock
promoters

10

Confirmation by RNAi Slight loss of G1, increase in G2/M indicating arrest in
G2/M

15

Line ID 238/44
Category Meiotic defects in testis: segregation defects, multi-stage defects (PI-02/18)
Reversion R
5 **Map Position** 70D

Rescue ID F8E

Rescue Sequence

10 GTTCAAACGCACTTTTAAGGTGGCCTATCGGCCCATCAGGAGCAACTTTGTTC
TGCTGCCGGATCAGTACTACGGCGTCGTTTCGACCTATGTAAGTGTCTAAAGG
TCTTCGCTCCATTCAGATTAGATACGCCAAAGATTAATCCGGTCAACCATTC
GATTAGGACACGGGCTGCCTGAGCTTGCAGTACAATGGTCGGACGCACTACG
CCTCCTGGGCGCCACAAAAGGGCGGGCGGCATTAAAGACACCGAGATTGG
GATCAATGCCAGAGCGGCCAAGGAGATCGGTAAGCCATTACTTAACGGCCGG
15 ATGTGCATCGGTTGCCAATGTGCCGTAATATTGGACTCCGGCCATTTGCCCCG
TACCTCGTACGCTAGCAGCACCCACTTACCCTTTCTTGCCGTATGTCTGCACGA
GAATGATCTGGTCAAAGTGTGCGCT

Other results same as for line 238/37

20

Line ID 428/5
Category Meiotic defects in testis: cytokinesis defects, segregation defects (seg-01/01)
25 **Reversion** ?
Map Position 70A

Rescue ID G4E

Rescue Sequence

30 GTTCAAACGCACTTTTAAGGTGGCCTATCGGCCCATCAGGGAGCAACTTTGTT
CTGCTGCCGGATCAGTACTACGGCGTCGTTTCGACCTATGTAAGTGTCTAAAG
GTCTTCGCTCCATTCAGATTAGATACGCCAAAGATTAATCCGGTCAACCATTC
TGATTAGGACACGGGCTGCCTGAGCTTGCAGTACAATGGTCGGACGCACTACG
CCTCCTGGGCGCCACAAAAGGGCGGGCGGCATTAAAGACACCGAGATTGG
35 GATCAATGCCAGAGCGGCCAAGGAGATCGGTAAGCCATTACTTAACGGCCGG
ATGTGCATCGGTTGCCAATGTGCCGTAATATTGGACTCCGGCCATCTGCCCCG
TACCTCGTACCTAGCAGCACCCACTTACCCTTTCTTGCCGTAGGTCTGCACGAA
AATGATCTGGTCAAGTGTGCGCTCATCGCTGACATTCTCAACCTGCGCA

40 Other results same as for line 238/37

Line ID 848/7
Category Mitotic defects in brain: cytokinesis defect. Meiotic defects in testis: cytokinesis defect. Multi-stage defects
Polyploidy, no overcondensation
5 **Reversion** PI-01/10
Map Position R
70D1-2

Rescue ID G1E

Rescue Sequence 1

GGCCACCTTAAAAGTGCGTTTGAACATTCTCGTCGTGGGCGTGTGCGAATTTA
GTACGCTCCTTCCTGGTTTAAATCATTTTCGCACTAAACTTCTGCTCTCAGCGG
AATTTACTTTTGCTTTATTAGAGATGGGAGCTCGCGCATCAGCTGAGCCGATA
CTTGCGCAACAGGTGATACAGCTGATTAGAGATGGCCCTTTTCAACTGTTCCC
15 AGCAGTGACCGCTGCCATAACCGTTTTTCAAATTTACGTGAGAACAGACATAA
AATAAATATTACAGCTCGTAGTAAATGTTATTCTATATTTAAAAGGAAATTGT
AATAGTTAAAACCTTGCAATGAATCAGTTACGTTCAAAAAAGGAAACACACTTT
AGTTTTTTGGCTAGTTTATTGGGTAAATAATTTTTATTTAAAATAGTTCGAGTG
TTCAATATAGTCATGTAAATGTGTACAGAAAGATCCGGCATTGATATTTAAT
20 ATATCGATTTCCCTTCACTTTCGCTCCTCGTATACCATGCTGGGGTCTTATCAA
ATTTATT

Rescue ID G1P

Rescue Sequence 2

AAGGTGGCCTATCGGCCCATCAGGAAGCAACTTTGTTCTGCTGCCGGATCAGT
25 ACTACGGCGTCGTTTCGACCTATGTAAGTGTCTAAAGGTCTTCGCTCCATTCAA
ATTAGATACGCCAAAGATTAATCCGGTCAACCATTTCTGATTAGGACACGGGCT
GCCTGAGCTTGCAGTACAATGGTCGGACGCACTACGCCTCCTGGGCGCCACAA
AAGGGCGGCGGCGGCATTAAAGACACCGAGATTGGGATCAATGCCACAGCGG
30 CCAAGGAGATCGGTAAGCCATTACTTAACGGCCGGATGTGCATCGGTTGCCAA
TGTGCCGTAATATTGGACTCCGGCCATCTGCCCCGTACCTCGTACGCTAGCAG
CACCCACTTACCCTTTCTTGCCGTAGGTCTGCACGAAAAATGATCTGGTCAAG
TGTGCGCCTCATCGCTGACGTTCTCAACCTGCACAGCGTCCACGTTACCCCCGT
CTCGTCCAA

Other results same as for line 238/37

Example 37 (Category 4)

Line ID 252/40
Category Meiotic defects in testis: segregation defects, abnormal spindles.
 5 (Ab-03/30)
Reversion R
Map Position 84E

Rescue ID A4B
 10 **Rescue Sequence 1**
 TACATGACTCTGCGATTTGACAAAAACAAAATTGAGTTTTGTCAAGAAAATCA
 ACTATTTTTCTGTGTTTAAAAAACCGAAGCCAACAAATCCGACCAAAATGCCT
 GCCGAAAACTTGGAGGAGCAGGGTCTGGAGAAGAACCCGAACCTGGAGCTGG
 CCCAGACGAAGTTCCTGCTTACCCTGGCGGAATACAAGCAGGATGCGGCATTG
 15 AAGGCGAAGCTTCTGGAGGCGATTTCGCACGGAGAATATGGCCCCGTGGGTAC
 GAGCACATCCTGCTCCGGAACCTCGGCTTGGACCCGTTAGACAAGGATCTTGCC
 TGGCGCCGAATTGAAGGAAAAACAATCGCGTTTAAGTTGGGAGCCA

Rescue ID A4E
 20 **Rescue Sequence 2**
 GTCATGTACTACCAGTGTGACCCCAAAGTTATCGATAAATTATACCGCATATT
 TTAACATTGCCAAAAATACCAGAGCGATGTCCATCAAGATAGCGACGAAATT
 AGAACAGTGCAATTGCCAATTGGGAATTTGTATTTTAATTTATTTTAAATTCT
 GAAAGTAATTTTAATTTAAAAAAAACCTTGAGAGCTGTCTAGAAAAGAACTTAT
 25 GTTTCATGATAACTTTGTGCGAAGAATTAAGAAATATTTAGTTGTAAAATAATT
 GTNTGAATCTATTTTTTTTCCAATAACACGACTTATATTTTTTTTTTAAATATTC
 CGAGCTAAATCCCAAGAAAGTTAAACTCCAATCTTGGGATTTTGAAGTGCCCC
 AGAAACTCCAAATTAACACTTCCTTTTTTAAATAATTGTTAAGACCCGTATCA
 CTTATGGTTATATACTGACCTCGAAAGGGCCACACTAAGGGGGGAGTTTGAAA
 30 ATTGATTTTCCTGATAAAAATTTTCGCTTGGAAAGCTACAGCATCGTCCACTGTC
 CATGTTTATATATCCTTATATTGCTTATAAATATAT

Genomic hit, Accession No. AC006494

35 **Associated ORF**
 Genscan: ORF1 predicted sequences >23:00:28|GENSCAN_predicted_peptide_2|389_aa
 MPAENLEEQGLEKNPNLELAQTKFLLTLAEYKQDAALKAKLLEAIRTENMAPWY
 EHICSELGWTVDKDLLARMKENNRVEVEQLDAAIEDAEKNLGEMEVREANLKKS
 EYLCRIGDKAAAETAFRKTYEKT VSLGHRLDIVFHLIRLGLFYLDHDLITRNDKA
 40 KYLIEEGGDWDRRNRLKVYQGVYSVAVRDFKAAATFFLDTVSTFTSYELMDYPT
 FVRYTVVYVAMIALPRNELRDKVIKGSEIQEVLHGLPDVKQFLFSLYNCQYENFYV
 HLAGVEKQLRLDYLIHPHYRYVREMRLGYTQLLESYRSLTLQYMAESFGVTVE
 YIDQELARFIAAGRLHAKVDRVGGIVETNRPDNKNWQYQATIKQGDLLLNRQKL
 SRVINI
 45 >23:00:28|GENSCAN_predicted_CDS_2|1170_bp

atgcctgccgaaaacttgaggagcagggctctggagaagaacccgaacctggagctggcccagacgaagttcctgcttacct
ggcgggaatacaagcaggatgcggcattgaaggcgaagcttctggaggcgattcgacggagaatatggccccgtgttacgag
cacatctgctcggaaactcggctggaccgtagacaaggatctgctggcggaatgaaggagaacaaccgcgtagaggtggagc
agctagatgcggcaatcgaggatgcggagaagaatctgggcgagatggaagtgcgcgaggcgaatcttaagaagtcagagta
5 cttgtgccgcacgcggcgacaaggctgccgcagagactgccttccgcaagacctacgagaagaccgtttccctgggtcaccgcct
ggacatcgtgttccatctgatccgcttgggactgttttaccctgaccacgatctcatcactcgcaacatcgacaaggccaagtatctg
atcgaggaaggcggcgattgggaccgacgcaaccgggtgaaggtctaccaggggtgtttactcgggtggcggtgcgtgacttcaag
gcggcgggccacgttcttcttgacaccgtaagcaccttcacctcatacgaactgatggactacccaccttcgtgcgttacaccgtt
tacgtggccatgattgccctgccgcgcaatgagctgcgcgacaaagtgatcaagggtccgaaatccaggaggtgctccatggc
10 ctgcccgcagtgaaacagttcctgttttctgtacaactgccaatatgagaacttctacgtacacctggccggcgtagagaagcaa
ttgcgcttggactacctcattcatccccactaccgctactacgtgcgcgagatgcgcattctgggctacaccagttgctggagtcg
tategctccctcaccctgcagtatatggccgagtcgttcggcgtaacagtggaatacattgaccaggagctggcacgcttcacgc
cgccggacggctgcatgccaagggtggatcgcggttggcggcattgtggagaccaatcgccctgacaacaagaactggcagttacc
aggcgaccatcaagcagggcgatctgctgctcaaccgcatccagaagttgagccgcgtgataaacatctaa
15

Drosophila Gene Hit BLASTN with rescue sequence 1 and TBLASTN with ORF1: 26S
proteasome regulatory complex subunit p42A (AF145308).
Human Homologue BLASTX with ESTand TBLASTN with ORF1: Hypothetical
protein KIAA0107 (D14663).
20 **Drosophila EST** several including GH17651 (AI387197)

Annotated Drosophila genome genomic segment AE003739
Annotated Drosophila genome Complete gene candidate CG5378 - Rpn7 19S
25 proteasome regulatory
particle, non-ATPase protein,
subunit S10aHuman
Homologue

Human homologue of Complete gene candidate gi7661914
8843E6684AE91ACD
30 [ref]NP_055629.1| KIAA0107
gene product [Homo sapiens]
(3.40E-149)

Putative function component of the 19S proteasome regulatory particle

Confirmation by RNAi Marked decrease in G1 and G2/M indicating fewer cycling
40 cells

Example 38 (Category 4)

Line ID 277/7
Category Mitotic defects in brain: anaphase defects
(weak, higher condensation, some polyploidy, fewer anaphases,
5 polyploids with monopolar spindles)
Reversion ?
Map Position 71B

Rescue ID B8E

10 Rescue Sequence
AGTCGGCGCATGCGGAGAGAGAATCGAAAGAGAAAGAGAAGCAAAGAGAGC
GACATACAGCAAAAACAATTCAAAAAGAACTGGTGAAGAATACGAAAATAAG
ATAATTTTTTTAAGGAAGTCGCGCTTTGATCCGTATCCGTTTTAGCGTCCAAGAT
TTATATCTTAAATCGGACCTATATTTTGAGGTACAGTGAAGCTTTGATGCGCCA
15 GTCTTATATGAGTTAAAGTTTTAACGATTGAAAGACACCCCTGAGCTGCTCAT
TATATTTCAATATTTATAAACAATCTTATATCAGAGCTTGAGAGACTTGCATGC
GCCACAAAATTCCAATTCCAATTCCAATTCCGGAATAATTTACAATAATCTC
AATTAACATACGTATTTTATGTTTCGTAATTTTTTAAAATTCCCAGATTCCCCAC
AATTGCCATAATAATCTCGATTATGTTATTATACTCTGAGAAGTAGGAGTGTG
20 TGCAAAGACCACAAACAAATCATTAGGGGGCGT

Annotated *Drosophila* genome genomic segment AE003584
Annotated *Drosophila* genome Complete gene candidate CG15383 – novel

25 Human homologue of Complete gene candidate none

Putative function No homologies to indicate function

Confirmation by RNAi Slightly increased G1 decreased G2/M indicating arrst in G1

Example 39 (Category 4)

| | | |
|----|---|--|
| 5 | Line ID | 284/4 |
| | Category | Mitotic defects in brain: anaphase defects (overcondensation, polyploidy (with overcondensation), few anaphases, metaphase with bipolar spindle) |
| | Meiotic Reversion | NR |
| 10 | Map Position | 89B |
| | Rescue ID | 2C6E |
| | Rescue Sequence | |
| 15 | | GTCTACCACTAGCTCTTTGTCTTCGCCTTCTAGTCTCTCTCATCTTGGCAGCCC |
| | | GTTCTAGTGCGCGTATTTTTAGTCGCAACACATTGCCCAATTCGCCAGCCGCTA |
| | | TTTGTGTCGTCCATTTGTTTCATTCATCGGGCTCTTTTCCGATTTTCAGTGGGTGG |
| 20 | | CATTTAACAATAATCCCTGCGTTCGCTGTCCACGTCCACATTACGATACGTTTA |
| | | GTGCACGGAAAGAAATAAGCGTGTGGTTTCATAATATTAGCTATTGAAAAAA |
| | | GTTCTTAAATTTAAGCCTCACTCGATTCTGATGCATGAAATATTATTGGATTGT |
| 25 | | AAATGAGCGTCATGTTTTGGTATACAAATCTCAAAGTAATTTAAAAATTCTCA |
| | | TCTTACCGTACCTTGAACCACTACCAATCATCTCAGTACAGCATTTTCAGCGAA |
| | | TTTCTCACTGTGCACTACAATGCCAGGCGGTACAAGCACCTGTATTTATTTATG |
| 30 | | GTCCGCTGCCGTAATCGACTGCAGTCGCCGCTTCCCTCTCTCTTTTGCTACCAA |
| | | CAACTTGGGGTAGGGCACCTGAACTAGTTTCAAACGGCGGCGGTCGGCCTTTT |
| | | CAGCTTTTTCGCATTTGCCATTTTCCCGCGG |
| 35 | Annotated <i>Drosophila</i> genome genomic segment | AE003711 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG4275 - mor transcription factor involved in chromatin remodelling |
| | Human homologue of Complete gene candidate | CG4275- 4507081 [ref]NP_003066.1[pSMARCC 2 SWI/SNF related, matrix associated, actin dependent regulator of chromatin, subfamily c, member 2(aa) |
| 40 | Putative function | Transcription factor, regulator of chromatin |
| | Confirmation by RNAi | Decrease in G1 and G2/M and increase in polyploidy |

Example 40 (Category 4)

| | |
|--|--|
| Line ID | 407/8 |
| Category | Meiotic defects in testis: cytokinesis defects |
| 5 Reversion | ? |
| Map Position | 64B1-2 |
| Rescue ID | A9E |
| Rescue Sequence | |
| 10 | GACTCACCCCTTTCACGCATTTTCATTGGAACGTTTGTTCGTTTATGCACACGC |
| | GTGTTGACACTTTTCATGAAACGCAGTGCGTGAAAAGTGCATCGCATAAACGC |
| | AATAAATGTTTGATGGATGCGTTCTGATGGCTTGAAGTCGCCTATTTGGCCGA |
| | TTTTCGCACGTCCACTCCCGACGGCAACAGAGTCCTGACTGAATCCCGGAGCG |
| | GAAGGAGTGTGGATAGCCAGGACTGCCAAAGGACACTGCGCACTTTTACTTTT |
| 15 | TCGAAAGCGAAAGCGAAAGTGGTGGGGCCCAGGCCAAAACAANCCCTTGAGT |
| | TGAAATTGGAAAAAAACCGGGACAGGGATGGGAGCCCAGCTCCAACAAACG |
| | GTTCCGGATTCCCTTGGGAAAGCCACGCCCTGCGCCTGGAAAAGGAATGCCCTC |
| | CACCTCATTTGTCCTCCGTTTTTGCCTATCTCTCCCCCAAATTTCCGTAAATG |
| | AAAAACAACCTTTGGGTTTTTGGTTTTTAACAATTTCTCCCCATTTGGTTTTNGGG |
| 20 | TTCCCTTTCCATTTTGGGAATTGGTTTTAATTAAAT |
| Genomic hit, Accession No. AC005814 64A6-64B6 | |
| Associated ORF | |
| 25 | Genscan ORF1 predicted sequences >22:57:22 GENSCAN_predicted_peptide_2 524_aa |
| | MGRRKDKPRVPEQDARICRAICLCQLTMVLSCVSIVYLSVAIYSPSLKAFKSGFEL |
| | DPVMCQTVDRQMPNNCPWASCGEWCLTKTSGFCPQIHIVRRNGTDIQLNNCTR |
| | VTNTSCAMIDLSRLNKFNCNNGTACNNIRGVFNCSNGHCKNMSEFFLCHHKADG |
| | LTVNSQKDNTKLNNGFFECHGVHCTKIKKPFSCDRYCSKITTTNVNTLIMHEDNLIA |
| 30 | ADCENAVAFNQARGSEHGVRIEPFEFWKEDDGNLLTNCATVTRESNDRITATDCI |
| | NGTLLEHDTLPAPFMNFTQFWAIYENSTRSVDPEQRYLPNQANLTIYSWKKLFINL |
| | EGCVNTLRGECKDFVARYGNDGDNNTAQSRYQCYYNKDSNVEFVVARYDLDK |
| | VYRELLVSLIVPIVLFVISSISLCITKSVKVGDDAKMRCVCAGDDSDNDGPFGPGL |
| | ANKQQDQMYDTHDDVVDLEHQAVDGQELSDHGLPLDNQELIGSTKSLIPSPVGE |
| 35 | SGTSDQIFDQDQEKATTCDVPEKPLVIL |
| >22:57:22 GENSCAN_predicted_CDS_2 1575_bp | |
| | atggggcggcgcaaggacaaaccgcgggtgattcccgaacaggatgcgcgcacatctgtcgcgccatctgcctgtgccagctgac |
| | catggtgctgtcctgcgtgtccatcgtctacctaagcgtggccatctactgcctccctaaaggcctcaagtccggcttcgagct |
| 40 | ggatcccgtcatgtgccagacggtggatgccagatgccaacaactgcccctgggcatcctgcggcgagtgggtgcctgacca |
| | agaccagtggctttgccccagatccactcaatagtgcgtcgcaacggcaccgatatccagctgaacaactgcaccagagtcac |
| | caacacatcgtgcgccatgattgacctgagtcggctgaacaagttcaattgaacaacggcaccgcctgcaacaatatcagagggc |
| | gtcttcaactgctccaatggacactgcaagaatatgtcggagttcttctgtgtcaccacaaaggccgatggacttacggtcaattcgc |
| | agaaggataacaccaagctgaatggattcttcgagtggtcacgggggtgactgcaccaagatcaagaagcccttcagctgcgateg |
| 45 | ctactgttccaagataacaactaccaatgtgaacacccttattatgcacgaggataatcttattgccgccgattgtgagaacgcagtg |
| | gctttcaaccaagcccaggatccgagcacgggtgtgcgtatgaacccttgagtttggaaagaggatgatggcaacctgctga |

ccaactgcgccacagtcacaagagagtcggacaatgcatacactgccacggactgcataaatggaacctcctggaacatgaca
ccttgcccgcctcccttcatgaacttcacccagttttgggccatctatgagaacagcaccagggtcggatcccagcagaggtac
ctgcccgaaccaggccaacctgacatctacagctggaagaaactgttcatcaacctggagggctgcgtgaacacactgcgtggg
gagtgaaggactttgtggctcgctatggcaacgatggcgataacaacaccgcccagtcacgctaccagtgtactataacaagg
5 actcgaatgtggagtttgtggtgcacgctacgatttggacaaggtttacagggagcttctagtctcgtgattgtgcccattgtgctc
tttgtgatctcatctatategttatgtatcatcaccaaatccgtcaagggtgggtgacgatgccaagatgcgctgtgtttgtgccggcga
tgattcagataatgatggcccctttggcccaggactagcaaacaagcagcaggatcagatgtacgatacagacgacgatgtagtt
gacctggagcaccaagcgggtggatggtaagaactatcgaccacggacttccgctggacaaccaagagctaatacggtagcac
caagtcgttgataccaatcagtcctcggtcgagaaatccggaactagtgtatcaaatctttgaccaggatcaggagaaagcaactacgt
10 gcgatgttcccagaaaaccactagtcatactataa

(corresponds to CG15003)

- 15

Annotated *Drosophila* genome genomic segment

Annotated *Drosophila* genome Complete gene candidate

AE003480

CG15003- novel unknown
- Human homologue of Complete gene candidate

none
- 20

Putative function

No homologies to suggest function
- Confirmation by RNAi

Only wild type profiles observed

Example 41 (Category 4)

| | | |
|---|--|---|
| 5 | Line ID | 422/28 |
| | Category | Meiotic defects in testis: segregation defects, multipolar spindles (Mul-02/22) |
| | Reversion | NR |
| | Map Position | 68E |
| 10 | Rescue ID | 2I4E |
| | Rescue Sequence | TCGTGGACCCTCAAAGNAACGGATTTCTCCAGTTTCTTCAAAGGGTTAATAAA CTTTTCGCACGTTTCGCATTTTTATGCTCAATCCGGTTACAAAATGCTGATAAA ACCACTTGAACCTACACGTTTCCGTACTGATAAGGGCTTTTCTTCTTATCTGACC TCTGGAATTCCGCGGAATTAATTCTTGAAGACGAAAGGGCCTCGTGATACGCC 15 TATTTTATAGGTTAATGTCATGATAATAATGGTTTCTTAGACGTCAGGTGGCA CTTTTCGGGGAAATGTGCGCGGAACCCCTATTTGTTTATTTTTCTAAATACATT CAAATATGTATCCGCTCATGAGACAATAACCCTGATAAATGCTTCAATAATAT TGAAAAAGGAAGAGTATGAGTATTCAACATTTCCGTGTCGCCCTTATTCCCTTT TTTGCGGCATTTTGCCTTCCTGTTTTTGTCTACCCAGAAACGCTGGTGAAAGTA 20 AAAGATGCTGAAGATCAGTTGGGTGCACGAGTGGGTACATCGAACTGGATCT CAACAGCGGTAAGATCCTTGAGAGTTTTTCGCCCCGAA |
| Genomic hit, Accession No. CSC:AC014962 | | |
| 25 | Annotated <i>Drosophila</i> genome genomic segment | AE003543 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG5684 (putative transcription factor, human homolog |
| 30 | Human homologue of Complete gene candidate | 1e-100 4758946 ref NP_004770.1 pPOP2 POP2 (yeast homolog) |
| | | >gi 4106061 gb AAD02685 35 (AF053318) CCR4-associated regulator of polymerase II transcription |
| 40 | Putative function | Transcription factor |
| | | |

Example 42 (Category 4)

| | | |
|--|--|--|
| 5 | Line ID | 422/5 |
| | Category | Meiotic defects in testis: segregation defects, abnormal spindles (Ab-04/26) |
| | Reversion | ? |
| | Map Position | 82D |
| 10 | Rescue ID | B9E |
| | Rescue Sequence 1 | ATTGGCTCTTGATGGACTACAACGCTACCAAAATGGGGCTTGAGTTGAATTAC CTGTTGGAAGACACAATGCCACCCACGATCAACAATTCGGCGGTAAACAGTG CCGCCGAAAAGCGACCCAGCGGCAAACGGGAGCGCAAGTAAGTGAACAGAT CCCTAAACAGACCCAGATACTCAGACTGATGTGTACCTTGCAGATCCGAGATC 15 ATTTGCCGCGTGAAGTATGGAAACAACCTGCCGGATATACCATTTGATCTGAA GTTTCTGCAGTACCCCTTCGACAGCCACCGCTTCGTGCAGTACAACCCAACGT CGCTAGAGCGTAACTTCAAGTATGACGTGCTGACGGAACACGATTTGGGTGTC ACGGTGGGACCTGATTAACCGGGAGCTCTATCAGGCCGACTCCATGACGCTGC TGGACCCGCCGATGAAAAACTGCTGGAGGAGGAGACTCTGACGCCACAGAC 20 TCTGTGCGTTCGCGCCAGCATTCGAGGACGGTGTGTCATGGTTGCGCAAATCCGA GT |
| 25 | Rescue ID | B9B |
| | Rescue Sequence 2 | GGCCAAATCTAGAAATCCTCAAATCTGCGCTTGGCAGTGTGACCGTACTTGAC CGGTACGATAATACCTCCGGTAAAAAAAATACTATATTTCCGGGGGACTCAAA TGCAACATCCTCATCGTATATAACACAACATCTATTTGAATTTTCATTTCCACAA CTAATATTATGGATAATGCTTTATTATCATTTTCCAAGTTAGCGATAAATCACC CCACAAGCTGAAAAATCAACGTTTAAAAACGATTGATATTTTTTTTAATACTTT 30 TTGGTTTTACTATTTGAATTTTTGTATACTTTTAGATTTTACTATTTTAATTTTC GTTTCTTCTAGCTGACTAACGGGTAAAAAAGGATCCGTCGACCTGCAGATCT CTAGAAGCTTGCGTTGCTGGCGTTTTTTTCCATAGGCTCCGCCCCCTGACGAGC ATCACAAAAATCGACGCTCAAGTCAGAGGTGGCGAAACCCGACAGGACTATA AAGATACCAGGCGTTTCCCCCTGGAAGCTCCCTCCGTGCGCTCTCCTGTTCCG 35 ACCTGCCGCTTACCGGATACCTGTCCGCCTTCT |
| Genomic hit, Accession No. AC008189 | | |
| Associated ORF | | |
| 40 | Genscan ORF1 predicted sequences >15:53:24 GENSCAN_predicted_peptide_3 211_aa | |
| | MRNANESSGKPKSKFVSNEFHAFSTICSIADSPAVSREKLKIDLAARKIPSASAPK GDSPLERFSRDLFTYLRVCRWGRFSAALFTAELLIVGGIVSSNRTSESSETGNPLA NEPDPLYMKLVDPMVAGESPKRMIKDQKDVGLKSTSSSEELRKLPKTRGRQKRFI RNPNYVKANEFYDKMLSSEYVSKRYKDLPPHPGFGADQPPA | |
| 45 | >15:53:24 GENSCAN_predicted_CDS_3 636_bp atgcgcaacgcaaataaatcgagcggttaaaccaaaatcgaaatttgaagcaacgaattccacgcattgtttcaacaattgttcaa | |

5 ttgccgattccccggctgtctctcgagaaaaattgaaaatcgatttagctgctcggaaaataccttcggcatcagcccccaaaggg
gattctccactcgagcgcttttcgcgggatctgttcacttacttgcgctccgtttgccgctggggtcgcttttcggcggcactgtttacc
gccgaattgttgatcgtgggtggcattgtgtctccaacagAACGTCAGAGTCTTCTGAAACTGGAAACCCACTTGCAAACGAGCCC
gatccattatatatgaaactggtggatcccatggtagcaggagaatcacctaaaaggatgattaaggatcagaaagatgtaggcctt
aatcaactagcagtagcgaagagctccgaaaattgccaaaacgcgaggctcgacagaagagattcattcggaatccaaactat
gtgaaagctaacgaattctatgataagatgtaagcagtgaatacgtagtaagcgggtataaggatcttccgccgcctcatccggga
tttgagcgggatcaaccgccagcatga

Corresponds to CG2503

- 10 **Annotated *Drosophila* genome genomic segment** AE003605
Annotated *Drosophila* genome Complete gene candidate CG2503 - novel possibly
RNA binding
- 15 **Human homologue of Complete gene candidate** 3287674 AC005239
(AC005239)
F23149_1(aa)
- Putative function** Possible RNA binding protein
- 20 **Confirmation by RNAi** Almost no G1 and broadened G2/M indicating arrest in
G2/M

Example 43 (Category 4)

| | | |
|----|--|--|
| | Line ID | 423/14 |
| 5 | Category | Meiotic defects in testis: cytokinesis defects, abnormal spindles (Ab-16/13) |
| | Reversion | R |
| | Map Position | 67B1-10 |
| | Rescue ID | E9E |
| 10 | Rescue Sequence | |
| | | GTTTGGCGTAAAAGCTTCGGCTGTGTTTGGTGCCCAAATTTTCCACTGCTTCT |
| | | CTTTTTGTGTATCTCTTATATCTTGTGCTTTTTTGTGTGTATGTTTTCTCGTTTC |
| | | TTTTTGCACACGCGCTTCGCGTTGCGGGCCAGCTGTTTTTGTGATAAGTGGT |
| | | TACGGTTTGTGTGTGCCAGCGGGTTTTCCTTAGTCGAACTGCTCGCGATGACTG |
| 15 | | ATTTTTCACAAGTGACTCAAAAACAGTCGATCGCCCTTTTAAGAAAACCCGCT |
| | | CAACGCACACAAAAGCGGTTTCTCTCTTTTTGTGCGTTCTCTCTTTTCACACTGA |
| | | CCACACGGAACGAAAAAATGATTACCGACCACACGGAAAGAAAAATTTATGT |
| | | CCAGACGAACTATTTTTGTCCAAGTAGCTGATTTGCATAACAATTTAAGCCA |
| | | CAAGAACTAGATTAAAATTTTACATTTAAATACATTATCAAATCCGAAATAT |
| 20 | | CAATAATTGTAATTTATCCTTACAAAATGTTA |
| | Genomic hit, Accession No. | CSC:AC020214 |
| | Drosophila EST | several including LP12306 (AI297868) |
| 25 | Annotated Drosophila genome genomic segment | AE003552 |
| | Annotated Drosophila genome Complete gene candidate | CG3967 - novel |
| | Human homologue of Complete gene candidate | none |
| 30 | Putative function | No homologies to indicate function |
| | Confirmation by RNAi | Only wild type profiles observed |

Example 44 (Category 4)

| | | |
|----|------------------------|--|
| | Line ID | 427/5 |
| | Category | Mitotic defects in brain: anaphase defects. Meiotic defects in testis: segregation defects, abnormal spindles (mitotic : Overcondensation, lagging chromosomes/less aligned metaphase with bipolar spindles, Meiotic: Ab-06/20) |
| 5 | Reversion | ? |
| | Map Position | 67B1-5 |
| 10 | Rescue ID | H4E |
| | Rescue Sequence | GTACAGCCTGAAGTGATCGTTGTTGTTTGAATCGGTGCTATCGGCGGTGCGC TTTGTGGGCATCTTTATCCAATTTGCTATGCGCGCTTGTCTTAAATTTTGAAC TGTATTCCAAGGGTTGCTTTGGCGGCTATCGATAGTATCGGCATGGTTACATTT 15 TAGTTTTATAACAAGAATTTTACAGGTATTTTGATTATCTGAGCTTAGTTTTAA GCAANAATATTATTGTTAAAAATTTAAAAAGTAAACAAGCTATTTTAAACAAGC ATTTAAACAAATAGTATTAATAATAATAAAAAATATATCGATATGTGTTGCAAAT GTTCGTTCCCTTAGTATTCTCTCATATTTATTTCAAATAAACTGTATAAAATAT CTGAAAAAGCGAACATATTTATTTAATTTTCATCGCAGATATCGATATCACAGC 20 GCTGCTATCGATGGTGTGTCTGTCGCAGTGCCTATCGCTTACCCTGCCATCGCT AACAAAAA |

Genomic hit, Accession No. CSC:AC020120

| | | | |
|----|-----------------------|---|--|
| 25 | Associated ORF | Genscan: ORF2 predicted sequences >22:06:07 GENSCAN_predicted_peptide_7 464_aa MPSEQHTNIKVAVRVRPYNVRELEQKQRSIIKVMDRSALLFDPDEEDDEFFFQGA KQPYRDITKRMNKKLTMEFDRVFDIDNSNQDLFEECTAPLVDAVLNGYNCSVVFV YGATGAGKTFTMLGSEAHPLTYLTMQDLFDKIQAQSDVRKFVGVSYLEVYNE HVMNLLTKSGPLKLREDNNGVVVSGLCCLTPIYSAEELLRMLMLGNSHRTQHPTD ANAESSRSHAIFQVHIRITERKTDTKRTVKLSMIDLAGSERAASKGIGVRFKEGAS INKSLALGNCINKLADGLKHIPYRDSNLTRILKDSLGGNCRTLMVANVSMSSLTY EDTYNTLKYASRAKKIRTTLKQNVLKSKMPTEFYVKKIDEVVAENERLKERNKA LEAKATQLERAGNSGFDPLELKTWYSKIDAVYAAARQLQEHVLGMRSKIKNINY RQTLKKELEEFRKLMCVDQRVCSQESF | |
|----|-----------------------|---|--|

>22:06:07|GENSCAN_predicted_CDS_7|1395_bp

| | | |
|----|---|--|
| 40 | atgccttcggaacagcatacgaatataaaagtggcggttcgcgtacggccgtataatgtccgtgaattggagcaaaaacagcgga gtattatcaaggtcatggatcgttcggcactgctgttcgatcccgacgaggaggacgatgagttcttcttcagggcgccaagcaac cgtaccgcgacatcaccaagcggatgaacaaaaagttgaccatggaattcgacagggtattcgatatagacaattccaaccagga tctgttcgaggagtgcacggcgccgctggtcgacgcggtgtaaatggatacaactgctcggtatttgatatggagccactggcg ccggaaaaacattcacaatgctgggcagcgaggctcatccgggtctgacctatcttaccatgcaagatctcttcgataagatccaa gcgcagagcgcagtgcgcaagttcgatgtgggggtatcctatctagaggtgtacaacgaacatgtgatgaatctgctaactaaac gggccctttaaacttcgagaggaacaatggcgtggtggtcagtggtctttgtctcacgccatctacagtgccgaggagctgc taagaatgctgatgctgggcaactctcatcgactcagcaccacacagatgccaatgcagagagttccaggtcacatgccatcttc caggtgcacattagatcacggagcgcgaagaccgacacaaaagaacggtcaaactatccatgatcgtctggcgggcagtgga gagggcgggcagttacgaaaggcattggagtgcgattcaaggaaggcgccagcatcaacaaaagtccttagcttgggaaattg | |
| 45 | | |

cataaacaagctagccgacggcttaaagcacatcccgtaccgcgactcgaacctgacacgcacatcctgaaggactcgttgggcgg
aaattgtcgacattgatgggtggccaatgtctcgatgagctcactgacctatgaagatacctacaacacccttaagtacgctagccg
agctaagaagatacgacgactctgaaacagaatgtcctcaagtccaagatgccaacccgagttctatgtgaagaagatcgacgag
gtggtagccgagaacgagcgactcaaagagcgcaacaaggcgctggaggccaaggccactcagttggagcgcgcccggcaat
5 agtggattcgatccgctggagcttaagacgtggtacagcaagatagacgctgtatatgcggccgcccggcagcttcaggagcac
gtccttggtagcgtagcaagatcaagaacatcaactaccggcgagacactgaaaaaagaactggaggagttcaggaagctgatgt
gtgtcgaccagcgagtggtgccaggagagtttttaa

Drosophila Gene Hit TBLASTN with ORF2: kinesin like protein 67a (U89264)
10 **Human Homologue** TBLASTN with ORF2: kinesin family member protein KIF3A
(AF041853)
Drosophila EST GH22018 (AI402731)

Annotated Drosophila genome genomic segment AE003552
15 **Annotated Drosophila genome Complete gene candidate** CG10923 Klp67a -
motor protein

Human homologue of Complete gene candidate 2e-58 4758646 kinesin family
protein 3B
20 >gi|3913958|sp|O15066|KF3B
_HUMAN KINESIN-LIKE
PROTEIN KIF3B and also
predicted peptide
ENSP00000166696
25 Gene:ENSG00000073652
Clone:AC015936
Contig:AC015936.00023
6.70E-91 (predicted kinesin?:
ENST00000166696)

Putative function motor protein involved in cytoskeleton organization and
biogenesis

35 **Confirmation by RNAi** Almost no G1 and broadened G2/M indicating arrest in
G2/M

Example 45 (Category 4)

| | | |
|----|---|---|
| | Line ID | 442/3 |
| | Category | Meiotic defects in testis: segregation defects. |
| 5 | Reversion | ? |
| | Map Position | 70D4-7 |
| | Rescue ID | H7E |
| | Rescue Sequence | |
| 10 | CGCAAGACTGTCTTCGATAGCAGAAGCGTTATTTTCGGAACATATCGTTTATCG | |
| | AAACTACAGTTGCTCAATACTGAACTGTCCAGCTTCGAGTAGCTGTGGCTCAA | |
| | ACCATTGTTGTCATCGATAAGCAATTGCAATTTTATTTGTTTGCTTAAAAAATT | |
| | AAAATATAAACTACGAGGATCAAATATACATACATATTCCCAATATGTTAGCG | |
| | AAAAAACATTTCTGCTCAAAAAAAGATGTTTAAATACAATGTAAGCTGTTCTA | |
| 15 | TGCATTGAACAAATTAACACATTGAGAGGTCGCTCTTATAAGTGCACATTTC | |
| | ATTTAAATATATTTTAATATATTCAAATATAGTATAGCAGTATAGCATTCAAAT | |
| | GTAAGTGTGGTTGGACTATCGCTGTAGTCCAAGAACTGCAGATAGTGTCATC | |
| | GCTAGCTTTGAAGCATCTCAAAGGAAAAAGGGCGATAATTCTGATTA | |
| 20 | Genomic hit, Accession No. CSC:AC017664 | |
| | Drosophila EST | CK02287 (AA141680) |
| | Annotated Drosophila genome genomic segment | AE003536 |
| 25 | Annotated Drosophila genome Complete gene candidate CG6650 - novel transacylase like | |
| | Human homologue of Complete gene candidate | none |
| 30 | Putative function | Transacylase |
| | Confirmation by RNAi | Marked increase in G1 indicating arrest in G1 |
| 35 | | |

Line ID 473/22
Category Meiotic defects in testis: no division
(no meiosis)
Reversion R
5 **Map Position** 70A1-5

Rescue ID 2B7E

Rescue Sequence 1

10 CGCAAGACTGTCTTCGATAGCAGAAGGCGTTATTTTCGGAACATATCGTTTTAT
CGAAACTACAGTTGCTCAATACTGAACTGTCCAGCTTCGAGTAGCTGTGGCTC
AAACCATTGTTGTCATCGATAAGCAATTGCAATTTTATTTGTTTGCTTAAAAAA
TTAAAATATAAACTACGAGGATCAAATATACATACATATTCCCAATATGTTAG
CGAAAAAACATTTCTGCTCAAAAAAAGATGTTTAAATACAATGTAAGCTGTTT
15 TATGCATTGAACAAATTAACACATTGAGAGGTCGCTCTTATAAGTGCACATTT
CAATTTAAATATATTTTAATATATTCAAATATAGTATAGCAGTATAGCATTCAA
ATGTAAGTGTGGTTGGACTATCGCTGTAGTCCAAGAACTGCAGATAGTGTCA
TCGCTAGCTTTGAAGCATCTCAAAGGAAAAAGGGCGATAATTCTGATAAGAA
AGTTGGCGTAGCCGGAAGGCGGATTGTCACATACAAAATAGTTTGGAAAGCC
CAAACGTAG

20 **Genomic hit, Accession No.** CSC:AC017664
Drosophila EST LD47104 (AI515336), SD03663 (AI532240)

For other results see line 442/3

25 **Line ID** 670/6
Category Meiotic defects in testis: segregation defects, abnormal spindles
(Ab-12/48)
30 **Reversion** ?
Map Position 70C

Rescue ID H7E

Rescue Sequence

35 CGCAAGACTGTCTTCGATAGCAGAAGGCGTTATTTTCGGAACATATCGTTTTATCG
AAACTACAGTTGCTCAATACTGAACTGTCCAGCTTCGAGTAGCTGTGGCTCAA
ACCATTGTTGTCATCGATAAGCAATTGCAATTTTATTTGTTTGCTTAAAAAATT
AAAATATAAACTACGAGGATCAAATATACATACATATTCCCAATATGTTAGCG
AAAAAACATTTCTGCTCAAAAAAAGATGTTTAAATACAATGTAAGCTGTTCTA
40 TGCATTGAACAAATTAACACATTGAGAGGTCGCTCTTATAAGTGCACATTTCA
ATTTAAATATATTTTAATATATTCAAATATAGTATAGCAGTATAGCATTCAAAT
GTAAGTGTGGTTGGACTATCGCTGTAGTCCAAGAACTGCAGATAGTGTGTCATC
GCTAGCTTTGAAGCATCTCAAAGGAAAAAGGGCGATAATTCTGATTA

45 **Genomic hit, Accession No.** CSC:AC017664
Drosophila EST CK02287 (AA141680)

For other results see line 442/3

Example 46 (Category 4)

| | | |
|----|--|--|
| 5 | Line ID | 460/20 |
| | Category | Meiotic defects in testis: segregation defects, multipolar spindles (mitotic: High polyploids, no diploids, higher mitotic index Meiotic: Mul-02/59) |
| | Reversion | NR |
| | Map Position | 78A1-4 |
| 10 | Rescue ID | 2B8E |
| 15 | Rescue Sequence | |
| | AGCTGGTCCAATTGGAAACGTTAGCTGCTCCAATGGGAGCAGCTGGCGCTCTC | |
| | TCTTCGATCGCGCTCGCTCTCATCCTCTCTCTTTAGCTTGTGCCACAGTAGCTG | |
| | CCGAAGGCAATTTTCATGTGCTCGTGTGTCGACCCCCACTCAGCCCCTTCTG | |
| | ATCGGAATCGGGGATTTCGGAATCGTGTAAGGCAGCCTTTGAAGGTCCCTTTTC | |
| | CAGGTGGCGGCCGTATCCTTAAAGTAAACATAGTTCAACTGACTTGGCAGCGC | |
| | TCCAAATGCGGTGACTTCTTGGCTATGTCATATATACCCCCACTCCCCTCCTGA | |
| | CTACCCTGCCACGCCCCACCGCCCACCGTCGGCGACGACAATTCCATTAAAAG | |
| | TTGTACGTTGTCACTTTTCGTTAACTTATCTGTGGAGCATGTTGTGCGATCGCA | |
| | TTTTTATTGTCGCCATTGTCTCTCGCTCTCTCCATCGCTCTTTCGCCTGGCTTCC | |
| 20 | CTACCCTGCCCACACAGGGAAGCCTACACACTCTTAAATCATGCACTTGGAAC | |
| 25 | AAAAAGTGCAAGCATTAAACTTTATTTAAACATTCAAGAGCCGCTTCTCTATT | |
| | TACCATTGAAAATTTAATTTAAAATAGAAGAGGCCTTTTCAGAATAATATAAT | |
| | ACCTTTAAG | |
| 30 | Genomic hit, Accession No. CSC:AC020460 | |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003592 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG10588 - novel gene with homology to proteases |
| 35 | Human homologue of Complete gene candidate | |
| | 2e-74 4505453 | |
| | ref[NP_002516.1 pNRD1 nardilysin (N-arginine dibasic convertase) >gi 2462488 emb CAA6369 | |
| 40 | Putative function | Novel protease |
| | Confirmation by RNAi | Marked increase in G1 indicating arrest in G1 |

Example 47 (Category 4)

Line ID 477/16
Category Meiotic defects in testis: segregation defect.
Reversion NR?
Map Position 90C5-10

Rescue ID C3E

Rescue Sequence 1

10 CTGTGGACGGTCGTCAATGCGTGAATATTCTTCTATGTGTAAGTGGTGTGCGT
 GTATGTAGATTTCTGGTTAAGAAAAGCCCCAAAAACCAAAGCGCCCCGCAAA
 ATATATATTGAGTCTTCTTGGCCCAACAACAAATCTGCCGCCGGACTTTCGCC
 GGAGGGCGAGTGAAAAATTCAGTTTCTCTCCTCTCGACGATGCACTTTGGAGG
 CTGTGTGAGTGTGTGTGCGAGTGAGTGCGTGTGTGTATACATATGCAAATGAT
 15 TGGATGTGAATCCTTGCATCATCATCTTCATAAACACTTGGCGAAAAAC
 CGCAGGAAAACGCAAGCAGCCGAACAAAAAAGAGAGCCTCTCAAGACAAC
 GGCAGCGGCCAAAAGTGAACGCGCAACAAACGCGGCCAAGCAGGCGCGGCA
 ATTATTTATAAATCTTAAGCCGTTAGCCCCCTCTCTCTCCCACTCACGAAAAG
 AAAATAAGTTAAACCAATTGGTGAAGATGATGCCCC

Rescue ID C3P

Rescue Sequence 2

GTCCACAGACTGGCTATATATACTAAAAACGAACTCGCGTGAGAAGACAGGG
 ACAGGGCAGCAAACCTCGGTATACGAACGGAACGAAATGAAACGATTCAAGTA
 25 GTAGTGTATGCAAGTCTTGTCTGTCTGCGCCTGGCGTTCTTTTCTCTTTT
 TCGATGGTTTTTCGCCAGGCTGGGCGCTGCCAAAACGCTGATACGGCGGCCAC
 AATCACACGCGGCTAATCGCCAGTTGGGCCCTGCACAGGCTGCACATACTTTT
 CACTATTAATGCGCTGTATTTCACTTATTTTTTCGAACAAATTCGCAGCATGACG
 AAGAAGCGAGCCTGTACAAGATTAGAGCGGGTAGCACGCACGATAGTATCGA
 30 TACGTACGAGTATTTGGCACTGCGATACATTATCGGTGCTCGTTTCGATAGCCC
 CCGATAGCTCTAGCACGAAATTTTATCGCTTTATCCATATTTTATACTATTTT
 ATTTATTGGACTTCAATGAATATTTAATTTACGTCTGGGTCGCTTTTTTAAATAT
 ATATGGTAATCAATAGCTGGCGAATTAGCGATATTTGAGTGTGACGCAAAAAT
 GAGTTGCATCGATATCGATTTCTCGCTACTCTGGGACGCCATCTTTATTGCGG

Genomic hit, Accession No. AC007810

Associated ORF

Genscan ORF1 predicted sequences >17:48:58|GENSCAN_predicted_peptide_2|349_aa

40 MSRILFILLLLIVTQLSELQAAAFSVRQNRFDVPLQTPAPLATSTESSKKPEKAT
 SGLLKKCLPCSDGIRCVPQIQCPAHVRMESHEKPQICDLPAGKFGYCCETGQNHT
 APKPETS PKERRSGFPTILSPAVLDEARRNFEHLMHGVAQIPVRRGFPDFAHGLVF
 HSTAKDDLHNFAISNSAIEQVM TTTQLFGKKEQVPVEDFITNNVPIKFTETPLAHHC
 QPPPVCGNIRSVYRSM DGTNNPEPQRS LWGAAGQPMERMLPPAYEDVPSASPA
 45 AICSYTYGLASRLAPVSVVNCCTFAWQLDWTTGMASGECVCVECMPAEWRLGQC
 PLLHEASSEMSRLLAKS

>17:48:58|GENSCAN_predicted_CDS_2|1050_bp

atgagtcgcattttattttttgttgctacttattgtgacgcaactgagcgcaggtgcaggcggcagcattttctgtgcgcaaaatcgtt
 ttgatgaagttcctgattgcagactcctgcacctctggccacttccactgaatcttctaagaaacccgaaaaagctaccagtgggtct
 5 gctgaaaaaatgccttccctgcagcgatggtataagatgcgtgccccaaatccagtgtcccggccacgttcgcatggaaagccat
 gaaaagcccccatttgcgatctcccggctggaaaattcggctactgctgcgagactggacagaatcacactgctcccaagccg
 gagacctctcccaaggagcgtcgatccggatttcccaccattctgtcaccgcagtttggatgaggcgcgtcgcaatttcgagca
 ctgatgcatggagttgcgcagattccggctgcgcctggctttccagattttgccatggcctggtttccactcgacggccaaggat
 gaccttcacaacttcgcatatcgaacagtgccattgaacaagtatgaccaccagttgttgggaagaaggagcaggtgcccg
 10 tagaagatttcacccaacaatgtgcccataagttcactgagactccgctggcacaccattgccaaccgccccagtttgcggc
 aatattcgggtctgtttatcgcagcatggacggcacttgcaataatccagaaccacagagatctctgtggggtgctgctgggtcaaccg
 atggagcgcgtgctgccccccgcctatgaagatgttcgctcagcttctcctgctgctatatgtagttatatctatggcatcgcatctcg
 tctggcgccgtgtttctgttgcaattgttgacatttgcattggcaattggattggaccactggaatggcgagcggggagtggtgtgt
 gtggaatgtatgccggcgagtggcgttgggccaatgcccggttgcctcatgaggcgtcgagtgaatgagccgcctcttggcta
 15 aaagctag

Drosophila Gene Hit rescue sequence: eyelid/osa (AF053091)

Human Homologue BLASTX with eyelid: KIAA1235 protein (AB033061) Brain
 protein 120 (AB001895)

20 **Drosophila EST** several including LD04852 (AA201670), LD24466

Annotated Drosophila genome genomic segment AE003718

Annotated Drosophila genome Complete gene candidate CG7467 - osa DNA binding
 putatively involved in DNA
 25 packaging

Human homologue of Complete gene candidate

CG7467 - 7e-25 2588991
 dbj|BAA23269| (AB001895)
 B120 [Homo sapiens] and
 30 O14497 SWI/SNF-
 RELATED, MATRIX-
 ASSOCIATED, ACTIN-
 DEPENDENT REGULATOR
 OF
 35 CHROMATIN SUBFAMILY
 F MEMBER 1 3e-67

40 **Putative function** transcriptional regulator

Confirmation by RNAi Only wild type profiles observed

Example 48 (Category 4)

| | |
|--|---|
| Line ID | 496/4 |
| Category | Meiotic defects in testis: segregation defects, abnormal spindles (meiotic: Ab-08/42) |
| Reversion | NR |
| Map Position | 65E4-7 |
| Rescue ID | 2C1E |
| Rescue Sequence | <p>GCACGATCGCTCTCTCTTGGCTCTCTCTATCACTCTCTGGACTCTCTCTCAGCA CCTTTGCTACCGTTTCGCAGAACAGGTGTATCGGTTTTCAAGGCAACTGTGATT TTTTAACTCAACATTCTATATCGAAAACCTTGTAAGAGGTCGGAATTTTTCTTGAG CGCCTAAAAGTGTGCAGTGAAATCATTTAATCCACTTCCGGTTGCAAAACAGG AATCACACATATGAAGTGATTAAAAATCATAGAAGGTTTGACACCTTCAAATA ATAAGAAAACAAAAATTTGTAAACTGTGATAATTTATTTAATTGAAATCTTAA TTTAATGGCCTACAAATCTGTTGAATATCCGTTGAATACACTTTTCCAGGGTGT GTCCTAGTCGGCTCCTCTTTGTTACCCAGTTTGCTGGTCTTCTTAGCCGCACA CCAGTTTATCGCTGTTTTGCCTTTGCGCTTTTCATTCATAAACAAAAACAATG TTATTGTTATTGCGGTGGCTGTAGATGTAAATGTAAATGTAGATGTAGAGGCT GCTTCTTGGG</p> |
| Genomic hit, Accession No. CSC:AC018039 | |
| Associated ORF | <p>Genscan ORF1 predicted sequences >19:35:36 GENSCAN_predicted_peptide_6 190_aa MVSEQFNAAAEKVSLTKRPSDDEFLQLYALFKQASVGDNDTAKPGLLDLKGKA KWEAWNKKQKGKSSEAAQQEYITFVEGLVAKYDNGMHKQEPNTCQARNATFR KSSECSLDQNTYTSSVTVIPAFHEGPKNSTASWPRIYRCYQRNQQAANCKWANTN SVCCKPHGKQSRRIIFAEFLAGHTVQILG</p> |
| <p>>19:35:36 GENSCAN_predicted_CDS_6 573_bp atggtttccgagcaattcaacgccgccgccgagaaggtgaagagcctgaccaagcgtcccagtgatgacgagttcctgcagctg tacgccctgttcaagcaggccagcgttggtgacaacgacaccgccaagccgggtctcctggacctgaagggcaaggccaagt ggaggcctggaacaagcagaagggcaagagcagcgaggccgccagcaggagtacatcacctttgtggagggcctggtggc caagtatgacaatggaatgcacaacaagaaccaaacacttgccaagcacgcaatgcgactcggttcggaaaagctcggaatg ctcgtggtatcagaatacgtatacgtccagtggtgacggttatacctgcattccacgaaggtccaaagaactcgacggcaagttggc caagaatttaccggtgctatcagcgggaaccaacaagcggccaactgcaagtgggcaaacacaaatagcgtttgcgggaaaccc cacggaaaacagagccgccgaatcatcttcgcagaatttctggccggccatacgggtgcagattcttgggtaa</p> | <p>Drosophila Gene Hit rescue sequence: melt (S144114) P element insertion site (AF174669), TBLASTN with ORF1: diazepam binding inhibitor (DBI) (U04823) and melted (AF205831)</p> |
| Annotated Drosophila genome genomic segment | AE003560 |
| Annotated Drosophila genome Complete gene candidate CG8624 melt - putative signal | |

| | | |
|----|---|---|
| | | transduction protein |
| | | CG8631 msl-3 - acyl-CoA- |
| | | binding |
| | | protein/diazepam binding |
| 5 | Human homologue of Complete gene candidate | inhibitor |
| | | CG8624- predicted gene |
| | | ENSP00000065899 |
| | | Gene:ENSG00000055889 |
| | | Clone:AC015904 |
| 10 | | Contig:AC015904.00014 |
| | | 1.70E-15 (unknown predicted |
| | | gene 1: ENST00000065899 |
| | | and AK022666 Homo sapiens |
| | | cDNA FLJ12604 fis 2e-29 |
| 15 | | |
| | | CG8631- gi5803104 |
| | | 0C85AE40FDF874CD |
| | | [ref]NP_006791.1 male- |
| 20 | | specific lethal-3 (Drosophila)- |
| | | like 1 [Homo sapiens] (1.70E- |
| | | 36) and Ensembl predicted |
| | | peptide ENSP0000006617 |
| | | Gene:ENSG0000005302 |
| | | Clone:AC004554 |
| 25 | | Contig:AC004554.00001 |
| | | 8.70E-19 (unknown predicted |
| | | gene 1: ENST0000006617 |
| 30 | | |
| | Putative function | CG8624: putative signal transduction protein |
| | | CG8631:acyl-CoA-binding protein/diazepam binding |
| | inhibitor | |
| 35 | | |
| | Confirmation by RNAi | CG8624: reduced G1 and G2/M Indicating fewer cycling |
| | | cells, CG8631: Increased G1 to G2/M ratio indicating arrest |
| | | in G1 |

Example 49 (Category 4)

| | | |
|----|---|---|
| 5 | Line ID | 523/19 |
| | Category | Female sterile. Meiotic defects in testis: cytokinesis defects, segregation defects (Mitotic: Less condensed chromosomes, nuclear bridges, Meiotic: Seg-01/02 |
| | Reversion | R |
| | Map Position | 75C1-4 |
| 10 | Rescue ID | 2B4E |
| 15 | Rescue Sequence | ACTGAGAGCATATTTGTGCACCAGAGGGCTGCATAACAACATTCTCTTTGTCC |
| | | ATTCGTTATACTTCGTATTCAGAATACATGTCATTCAGTTGGTCCCGTTCTTTTT |
| | | GCGTTCACTTCGTATATATTCGGCGATCGAAATGAACTAACTGAATGTGTTCA |
| | | AAGAATGAATGAAGCCAATGAATTTTCAATAGTAATTCAGAGTGCTTAAAATT |
| | | CTTCATGTTGTCATTGAGTAAAATGAGTTCGGACAGCGCGAAGGTAAGTCGAA |
| | | GTTTGTGTTTTATTATGTTTATTTGTATTATTATGTACACTAGTCGGCATACTTT |
| | | TGCGTGCGTCTTATACGTGTGCGTCTTATTTAACAATATTGTAAAATAAAATAT |
| | | ATAAATTATTTGTTATATGCGTAGGGGCCTTTATTTTGTGTATTGATAGTCTTTT |
| | | GTCATAGATATCATTATTCTGACAAGATTTGAACTTTTCAAGTTATTGCCTCTC |
| | | GTTATTCAATTCCTAGCTGGTCTTACGTTACGCGATATTCCTAAAATATCCTA |
| 20 | | AAATCGCACAAAACAGTCACGCCACACTTTTGAAAAACGTGGTAATATTTTT |
| | | CATACTTGCATTAAGTCTGG |
| 25 | Genomic hit, Accession No. | AC007691 |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003520 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG4306 – novel |
| 30 | Human homologue of Complete gene candidate | 4e-25 3242764 (AC005154) similar to protein U28928 (PID:g861306) [Homo sapiens] |
| 35 | Putative function | No homologies to indicate function |
| | Confirmation by RNAi | Only wild type profile observed |

Example 50 (Category 4)

| | | |
|----|------------------------|---|
| | Line ID | 666/19 |
| 5 | Category | Mitotic defects in brain: anaphase defects (weak, overcondensation, aneuploidy, lagging chromosomes, metaphase with bipolar spindle) |
| | Reversion | NR |
| | Map Position | 64E1-5 |
| 10 | Rescue ID | I9E |
| | Rescue Sequence | CCCTCGTCTACGTCGAAATTCTGGATGCTTCTCGGATTTAGGGTTGTATCTCGA AAACGTTTGACTGCGAATGTCAATATCGATATGCTAACCGATAGCTGTCGATG TTTTAAACACAGTGCACTGTTTTTAAATCGCTCCCCATTTATATATATTTGTGC 15 NTGCTTTTGGCGGTNNTTTTCTTTATATGCTTCTTATGCTTTTACGATTATTATT AGCGCTTATTTGATTGCAAATGCCAAGGAAAGCGTGACTGTGATGGCGAAATG CGGAAAGTACTCCTTAAATCTCATATATCGCATAAAACTATCGGTTCTGGAAT GTTTCGTGTAAGTCTGCGAAGATAGAGATCGATCTATTTTGAGGATACATTTG TTAATATTATAAGGGATTCTTCTACAGGGGTCAGATTGCTTAAAAACACACAG 20 AANAATAAACAAAATATTTCTTTGAAATATTGAAATATTTGAAATANAAAAAA CGTATTGACGAGGTAAGCATATTGAAAAAGATAGGAAGGTGATGGAGAAAGT GCACTTATATTGGTCACCAAAGAGCTTATAATCAAAGATCAATAGATATAAA TATCTTTATATGATATAAAATATAATACATATAATATAATATCATATACAATG GATAAATTGCAAGTGGCAAAATGAATTCCGCGGAATTAATTCTGAANCGAAA 25 GGGCCT |

Genomic hit, Accession No. CSC:AC014815**Associated ORF**

| | | |
|----|--|--|
| 30 | Genscan ORF1 predicted sequences >17:46:43 GENSCAN_predicted_peptide_1 334_aa MGKDFYKILGLERKASDDEIKKAYRKLALKYHPDKNKSPQAEERFKEIAEAYEVL SDKKKRDIIFDNYGEDGLKGGQPGPDGGGQPGAYTYQFHGDPRATFAQFFGSSDP FGAFFTGGDNMFSGGQGGNTNEIFWNIGGDDMFAFNAQAPSRKRQQDPPIEHDLF VSLEEVDKGCIIKMKISRMATGSNGPYKEEKVLRITVKPGWKAGTKITFPQEGDS 35 APNKPADIVFIIRDKPHSLFKREGIDLKYTAQISLKQALCGALVSVPTLQGSRIQV NPNHEIIKPTTTRRINGLGLPVPKEPSRRGDLIVSFDIKFPDTLAPSLQNQLSELLPN | |
|----|--|--|

>17:46:43|GENSCAN_predicted_CDS_1|1005_bp

| | | |
|----|--|--|
| 40 | atgggcaaagacttctacaagattctgggcctcgagcgcaaggccagcgacgatgagatcaagaaggcctaccgcaaactggc actcaaataccatcccgacaagaacaagagcccacaggcggaggagcgcttcaaggagatcgccgaggcgtaggagtgctg tcggacaaaaagaagcgcgacatcttcgacaattacggtaggagtgattgaaggcgaggacagccgggaccagatggcggcg gtcagccgggagcgtagacttaccagttccacggcgatccgaggggccacatttgcccagttcttggatcgtaggagtcggttggc gcgttcttaccggcgggcgataacatgttagtgggcggtcaggcgggcaataccaacgagatcttctggaacattggcggcgacg atatgtttgcctttaatgccagggcaccagtcgcaagcgccagcaggatccgcccacgagcatgatctgttcgtgctgctggag 45 gaagtggacaagggatgcatcaagaagatgaaaatctcacgcatggccaccggaagcaatgggcccgtacaaggaggagaag gtgctgaggatcacagtgaagccgggctggaaggccggtaccaagattacctcccccaagagggtgattcggcgccaaacaa | |
|----|--|--|

gacgccagctgacatcgtcttcattcgcgacaaaccgcattcgtgttcaaacgcgaggggaatcgatctaaagtatacagccc
agatcagtcgaagcaggccttgtgcggagcactggtagtggtgcccacgctgcagggcagcaggatacaggtgaatccgaacc
acgagatcatcaagcccaccacaacgcgcccggatcaacggactgggtctgccgggtgcccaggagccatcgaggcgcggcg
atctgacgtctccttcgacattaagttcccgacacactggcaccagctctgcagaatcagctgtccgagctgctgcccactag

5

Drosophila Gene Hit rescue sequence: fasciclin I (FasI) (M32311) TBLASTN with
ORF1: DnaJ homolog (DROJ1) (U34904)

Human Homologue TBLASTN with ORF1: DnaJ-like heat shock protein 40 (HLJ1)
(U40992.2)

10

Annotated Drosophila genome genomic segment AE003565

Annotated Drosophila genome Complete gene candidate CG10578 - DnaJ-1 a
chaperone putatively involved
in protein folding. Stimulates
activity of HSP70

15

Human homologue of Complete gene candidate 8e-94 1706473 P25685
DNJ1_HUMAN DNAJ
PROTEIN HOMOLOG 1
(HDJ-1) (HEAT SHOCK
PROTEIN 40) (HSP40)

20

Putative function Chaperone involved in protein folding

25

Confirmation by RNAi Almost no G1 peak, increased G2/M indicating G2/M arrest

Example 51 (Category 4)

Line ID 714/11
Category Meiotic defects in testis: cytokinesis defects, abnormal spindles
5 (Ab-01/04)
Reversion ?
Map Position 66A10-15

Rescue ID 2A4E

10 Rescue Sequence
AACCAGAACGAACTCCAATGCAGTTTCATTTTGTTCAGTTTAATCATTAAACA
AAGAACTGCGCAACCGATCGCAACTAGCTCGTGGACTCTTGTTCTCCCAATAA
TTGGTATGTTTTCCATTTTTCGTTAACATGGAAAATGTGTGAAAAGCTTTTTCC
CCCTCCAAAAGAAGCGTACTGAACTAAGCTTTCGGTGGTTAGTAATAGTAGTC
15 GTTATATCTTATTTTTCTTATTTACGTGCAGCTGCAATCATTGGCTGCGTCACTT
TGGCGTCAGCTATAAACTGGTGGATCAACTCGGCGGCCTCCAAAAGCTGCGCA
TCTGCTCCAGACACTTTAGCCAACGCCAGGAGATGGCCAAAACCCGCATCAA
GATGACGCCGCTGCGCAAGTCCTCGTCCTCCAAGGGCATTGTGCTACCCATTA
ATGCCGCTGGAGGGTTCGGTCATTGCAGGCGCCTTAGCACGAGGAGGAGGTGC
20 A

Genomic hit, Accession No. AC012390

Associated ORF

25 Genscan ORF1 predicted sequences >19:47:45|GENSCAN_predicted_peptide_2|711_aa
MRSHQAVGNLLLADEALPAVQSASVYVWMAEQPLSPGQSYDIKIADSPSVSS
KSITDNGADVQWFAFEHSQYYQGQVQQMFLSALERIDSEFLITLIKRCPYHVDLSLVQ
LSEVCKMTEDFSLASELLERALLLLESSLHINFSLTSGNCRLDYRRQENRSFYIVLF
KHAQYLEERACSRTAFEISKLLLSLQPDTPDLAMILPNQPDQCTGNMTQLQQAGK
30 IRKRSEKQFPIGTEPRGTDALRFTLQTLASAGRDITWNIKRLQGSRVTGAAQGYLI
DKKTAVQYKITIAHLKDPNIDQLFDSSGDGKADLHGSTPDWGCQAMMADAISR
YKEGNPVFYITWTPYWVSNELKPGKDVVWLQVPFSALPGDKNADTKLPNAGGI
EGLIADEEVQVLDALCDAPCVGVSHSCRLLDGNRRGNNELRLFIPGKSQFGVADG
CADKQSVMEYHAAKTGHTKFSESEEEKKALTEEEKKAQLALIEEKLKQKRIEREE
35 REKIEALQREKNRIKSGKDMTEAKRRMEELEMKKIVEQRKREKDEEKAARDRVK
AQIEADKAARKAREQKELGNAEPAPSVSSTTVSSPPAGVKSPPRDYTETRIQGASA
ILAAAAPYYQPPAVPQDVQPDRPIGYGAFGVVCGSHISGWHCSAGHYEDGNENFE
CLKTFSTSDRIGCEWRWAAATVLAATCISPNGRCGHYKRVRRIKTNITTT

40 >19:47:45|GENSCAN_predicted_CDS_2|2136_bp
atgagatcgcatcaagccgttggaatctgctgctggcggcgagacgaagcgttacggcggtgcagagcgcgctcggtgtatgtg
gtatggatggcggaacagccgctttctcagggcagagttacgacatcaaaattgccgactctccatcggtgtcctccaagtctatc
acagataatggagcggacgttcaatggtttgaccttgagcatagccaatactaccaggagtgagcaaatgttcctttctgctctcg
agcgcatlgactcggaatttctgacacacttatacaacgctgccccatcatgtcgactccttgggtcaactcagcgaagtatgcaa
45 gatgaccgaagacttttcttgacctccgaactgcttgagcgcgcccttctcttctggaatcgctgctgcacatcaacttcagttga
cgtcgggcaactgccgactggactaccggagacaggaaaaccgatccttctacatcgctgctgtcaagcacgcgcagtlacctgg

aggaacgagcttgagccgcaccgccttcgagatctccaaactgctcctgagcttcagccagacacagatcctttgccatgatt
 ctaccaaatacagccggatcaatgtaccggcaatatgacgcagctgcagcaggcgggcaaatccgtaagcgctcagaaaagca
 gttccgatacgggtactgaaccgcgcgggtactgacgcgttgccgttcaccctgcagacactggcgtctgccggtcgcgacatcacct
 ggaatataaagcgctctgcaagggtcccgtgttaccggcgccggcccagggttacctcatcgataagaaaaccgccgtccagtacaa
 5 aatcaccatcatcgctcatctgaaagatccgaatatcgaccaactgttcgattcaagcggcgacggaaaagcggatttacacggta
 gtaccccagactggggctgccaagctatgatggccgacgccatcagtcgtacaaagaggggcaacccgggtgtttattacacctg
 gacgccgtactgggtgagtaacgaactgaagccgggcaagatgtcgtctggttgaggtgccgttctccgactgccgggcca
 taaaaacgccgataccaaactgccgaatgccggtggcatcgaaggcctcatcgccgatgaagaagtcaggtcctcgatgccct
 ttgtgatgcgccgtgtgttggtgtctcccactcgtgccgactccttgatggcaatcgccgagggaataatgaactgcggctcttatt
 10 cccggcaaatccagtttgagtagctgatggatgtgcagacaagcagagtgttatggagtaccatgccgcaaaaccgggtcac
 accaaattctccgaatcggaggaggaaaagaaggcgctcaccgaggaggagaagaaggcccagctggccctcatcgaggag
 aagctcaagcagaaacgcacatgaacgcgagggagcgcgagaaaatcgaagccctgcagcgggaaaagaatcgcatcaagtcc
 ggcaaggacatgaccgaggccaagcggcgcatggaggagtggagatgaagaagatcgttgagcagcgaagcgcgaaaa
 ggacgaggagaaggcggcccgatcgggtaaaggctcaattgaggcggacaaggcagcacgcaaggctagagaacaaa
 15 aggaattgggcaacgcagagccagctccatccgtgagctccaccacagtttcgtaccaccggccggtgtgaaatcctccgccg
 gagactacaccgaaacccgcacatccaggggcgccagcgcgaatcttgccgcagcgggtccctactatcaaccgccggtgttccc
 caggatgttcagccggatcgtcctatcggctatggagcattcggagttgtctgcggttcccacatcagcgggtggcattgttctgcg
 gggcattatgaagatggtaataaaaatttcgagtgcctcaagacatttcgacttctgaccgcattggctgcgaatggagatgggcg
 gcagcaactgttcttgcgcaacctgcattagcccggaacggccgttgccgggcattataaacgcgtacgtcgtcgcattaaaacaaa
 20 cataacaactacgtga

Drosophila Gene Hit rescue sequence and BLASTX with EST: BIP1 (Y14998),
BLASTX with genomic sequence matches BIP.

Human Homologue BLASTX with BIP1: alanine:glyoxylate aminotransferase
25 (X53414) ?

Drosophila EST GM04749 (AA695904), GM13608 (AA803601)

30 **Annotated Drosophila genome genomic segment** AE003556
Annotated Drosophila genome Complete gene candidate CG7574 - bip1 unknown
 function

CG13681 – unknown

35 **Human homologue of Complete gene candidate** none

Putative function no homologies to indicate functions, Drosophila Bip1 interacts with
transcriptional activator Bric-a-brac which is required for ovariole
formation

40 **Confirmation by RNAi** Both show reduction in G1 and G2/M indicating fewer
cycling cells

Example 52 (Category 4)

| | | |
|----|--|---|
| | Line ID | 763/4 |
| 5 | Category | Meiotic defects in testis: segregation defects (overcondensation, fewer anaphases) |
| | Reversion | R |
| | Map Position | 90F |
| | Rescue ID | 2F5E-1 |
| 10 | Rescue Sequence | CGGCAATGTCTGCGCCCCCAATCTGAACTTGCCTCGCCCTCTCCGCCCCCTGATC TCATCTCCTCTTCAAACCCCTGCTCCCCTTTTCTGCACACATTAACGTCAGCCT TTAAGTGTGCTTTCTCAGGTGCTGCCCCCTGCGCCCACCATCCCCCGCTCCATG CTCTTTCCATCTTGCGCTCTCTGCGTTCTATCTACATTTTTTTTCGAGGTCGCGCG 15 CTGCTTTTTCCGTTGATGTTCTCGTTCTCGTCAATGTCGCAATATGCGCAAAAGGC AGACAAAAAAAAAAATGAGTGGAAGTACATACATACCGGTGATTGATGGG CGGTGGGTGGCGGTGGTGTAGGNGTGGTTTG |
| | Genomic hit, Accession No. | AC006495 |
| 20 | Associated ORF | Genscan ORF1 predicted sequences >22:47:02 GENSCAN_predicted_peptide_3 283_aa MTERENNVYKAKLAEQAERYDEMVEAMKKVASMDVELTVEERNLLSVAYKNVI GARRASWRIITSIEQKEENKGAEKLEMIKTYRGQVEKELRDICSDILNVLEKHLIP 25 CATSGESKVFYKMKGDYHRYLAEFATGSDRKDAAENSLIAYKAASDIAMNDLP PTHPIRLGLALNFSVFYYEILNSPDRACRLAKAAFDDAIAELDTLSEESYKDSTLIM QLLRDNLTLWTSDMQAEEIPIKLPDRQSKTTLIFSPRSQVNPILHKNNTIIGRVIC SVFA |
| 30 | >22:47:02 GENSCAN_predicted_CDS_3 852_bp | atgactgagcgcgagaacaatgtgtacaaggcaaagctggccgaacaggccgagcgtacgacgaaatggtggaggccatga agaaggtgcctccatggacgtagagctgaccgtcgaggagcgaaatctgctgtcgggtggcgtagaagaatgtgattggagcac gccgtgcctcgtggcgcatcatcacctgatcgacagaaggaggagaacaagggggccgaggagaaattggagatgatcaa aacctaccgcggacaggtggagaaggagctgcgcgacatctgctcgatatactgaacgtgctcgagaagcatctcattccatg 35 cgccacatccggcgaaagcaaagtattctactataagatgaagggcgactaccatcgctacctggccgaattcgccaccgggtcc gaccgcaaggatgcggcgagagaactcgctgattgcctacaaggcgccagcgatattgccatgaacgatctgccaccaacaca ccccatccgttgggcttggcattgaacttctcgggtgttctactatgagattctcaactcgccggaccgcgcttggcgcttggcgaaa gccgcttccgatgatgccattgccgagttggatactgagcgaagagagctacaaagactcgacactcatcatgcagctgctgc gagacaacctcacattatggacgtccgatatgcaggcagaagagattccgattccaaaactccccgacagacagtccaaaacca 40 cattgatttttagccccgaagtcaagtaaacccaaagattctccacaagaacaacaccatcatcggcagagttatctgtagcgtgtt tgcgtga |
| | Drosophila Gene Hit | rescue sequence: 14-3-3 epsilon isoform gene (U84898) TBLASTN with ORF1: 14-3-3 . |
| 45 | Human Homologue | TBLASTN with ORF1 and BLASTX with 14-3-3: epsilon isoform 14-3-3 protein (U43430.1) |

| | | |
|----|---|---|
| | Annotated <i>Drosophila</i> genome genomic segment | AE003721 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG8045 complex gene |
| 5 | | appears to encode 3 things : Transcript: CT24102 unknown Transcript CT24072: transcription factor RNA polymerase II transcription factor , |
| 10 | | Transcript: CT24092: diacylglycerol- activated/phosholipid dependent protein kinase C inhibitor /14-3-3 protein epsilon (suppresspr of ras) |
| 15 | | |
| | Human homologue of Complete gene candidate | CT24092: e-119 NP_006752.1 tyrosine 3- monooxygenase/tryptophan 5- monooxygenase activation protein, epsilon polypeptide; 14-3-3 epsilon [Homo sapiens |
| 20 | | |
| 25 | Putative function | transcription factor, or 14-3-3 proteins which associate with cdc25 phosphatases |
| | Confirmation by RNAi | CT24102: wild type profile only, CT24072: Loss of G1 peak CT24092: Increase of G1 peak |

Example 53 (Category 4)

| | | |
|----|-----------------------------------|--|
| | Line ID | 951/8 |
| | Category | Mitotic defects in brain: (some overcondensation, anaphase bridge, metaphase with swollen chromosome and bipolar spindle) |
| 5 | Reversion | NR |
| | Map Position | 73D |
| 10 | Rescue ID | 2E8S |
| | Rescue Sequence | GTATAACAAGATCCCGAGACACCGGTCAGTTGGTGCTACACGCTCTTGGAGA GCGCTGTGTTTGTTCGGTTCAGCGATTAGCGATAGTTTTGTTCGAGCCGGTTGT GTTAACTTGGCTAGCTTCGGGTTTATTGTGACACTTTCCCCAAATCGATCGTTT 15 GCGAAGCGTGCATAGCGGAACATACATACATAGATAACCAGCGTGTCTGGGT GTTTCATGAAAAAGAGTGCGTGATATGGGATTCGATATGGCAACACGCTTTATG GATATACTAAAGCTGACCTTTAAGTGAGTTTTCCCGAGTCAGTGTCCGCTTCTTG CTCTTGCGGAGCGTTAAACGGTTTTCTGTGTTTTGAGGTCTCGCGTCTTGGTTT TGCAACAGCTTCTGCCCAGCATGCACACATACGTGTGCACTGGGAAAATAGTG 20 TTGCAGAAGTGCTTGATTTATAAATATTACAAAAAATGTGATGAAACACTTTT TATTTTCTTCAAAAAATCAAGAATAAATTAACACTATCCTGCTCTTAAACAT GGAGATTAATTCAATTTTAATTAAAAAATAATTTTTTTTACAATTTATGATTTA TGAATTTATGCACTCCTTGAAACTATTAAAGACTCAACAGTGA |
| 25 | Genomic hit, Accession No. | CSC:AC015272 |
| | Associated ORF | Genscan ORF1 predicted sequences |
| 30 | | >23:03:05 GENSCAN_predicted_peptide_1 602_aaMGFDMATRFMDILKLTFKPFKTN YTEEKYFNDKLRSSKNIERRYILDVGRGPTAVTYNPIWVISFKYEQRKLSTAIYSV IKTKSGPVRGVKRNTIWGGSYFSFEKIPFAKPPVGDRLFKAPEAVEPWDQELDCTS PADKPLQTHMFFRKYAGSEDCLYLN VYVKDLQPDKLRPVMVWIYGGGYQVGEA SRGLDVVIVTVAYRLGALGFLSLDDPQLNVPGNAGLKDQIMALRWVQQNIEAFG GDSNNITLFGESAGGASTHFLALSPQTEGLIHKAIVMSGSVLCPWTQPPRNNWAY 35 RLAQKLGYTGDNDKKAIFEFLRSMMSGGEIVKATATVLSNDEKHHRILFAFGPVVE PYTTEHTVVAKQPHELMQNSWSHRIPMMFGGTSFEGLLFYPEVSRRPATLDEVGN CKNLLPSDLGLNLDPKLRNYGLQLKKAYFGDEPCNQANMMKFLELCSYREFW HPIYRAALNRVRQSSAPTYLYRFDHDSKLCNAIRIVLCGHQMRGVCHGDDLCYIF HSMLSHQSAPDSPEHKVITGMVDVWTSFAAHGDPNCESIKSLKFAPIENVTNFKC 40 LNIGDQFEVMALPELQKIEPVWNSFYAPNKL |
| 45 | | >23:03:05 GENSCAN_predicted_CDS_1 1809_bp atgggattcgatatggcaacacgcttatggatataactaaagctgaccttaagccatttaaaacgaactacactgaagaaaagtattt caatgacaaactcagatcttcgaaaaatattgaaaggcggtatatcttgatgttggtttcgcgaccacagcagtcacgtacaat ccaatctgggtaataagcttcaagtacgagcagcgcaaattgtcaacagcaatatattccgtcataaagacgaaatcaggtcctgtg cggggagtggaagagaaacacaatctggggaggaagctacttcagtttcgagaagatacccttcgaaagcctccggtgggagat |

165

ctgcgcttcaaggccccggaagcagtgaggccatgggatcaggaattggattgcacttcgccggcagacaagcccccttcagaca
 cacatgttttcagaaaatacgcgggctcagaggactgcctctacttaaatgtgtatgtcaaagatctgcagccggataaactgcgtc
 ccgtgatggtttggatctacggaggaggctatcaagttggcgaagcttctcgaggattggatgtggtcatagtcaccgttgcttatcg
 actgggtgccttgggcttcctcagcctggatgatcccaactaaacgttcccggaaatgcaggtctcaaggatcaaatcatggccc
 5 tgcgatgggtgcaaaaaacatcgaagcattcggcgggtgattccaacaatattacactcttggcgaaagtgccggcggagcctc
 gaccacttccttgactaagtccccaaactgaaggcttatccacaaagctatcgttatgtcgggcagtgtttgtgcccctggacg
 caaccaccgagaaataattgggcttataggctggcccaaaaattgggatacaccgggtgacaataaggacaaggcgatctttgagt
 ttctgcgatcaatgagtggcggggagattgtcaaggccaccgcaacagttctcagcaacgatgaaaagcatcatcggtatcctttc
 gccttcggacctgtcgtagaaccatatactaccgagcacactgtggtcgtlaaacaaccgcatgaactgatgcagaatagctgga
 10 gtcacaggataccatgatgtttggaggcacgagcttcgagggttctattctatccagagggttcaaggcggccagcaaccctc
 gatgaggtgggtaactgcaagaatctgctaccgagcgcgtcgggtcttaacctagatcccaaactgcgtgagaactacggcttgca
 actgaagaaggcgtatttcggcgacgaacctgttaaccaggcaaacatgatgaagtttctcgagctatgctcatatcgagagttctg
 gcacctatatacagggcagcttgaaccgtgtccggcaatccagcgcacccacgtatctgtatcgattcgatcacgattccaaact
 gtgcaacgccattaggattgtactttgcggccatcagatgcgaggtgtttgcatggtgacgatctgtgctatatttccacagcatgtt
 15 gtcgcatcaatccgctcccgttctccggaacacaagggtataaccgggaatggtcgacgtttggacgagtttcgcagcccacgga
 gatcccaactgcgaaagtataaaatcactcaagtttgacccatcgaaaacgtaaccaactttaagtgtctcaatattggggatcagt
 ttgaagtcatggcgttccagaattgcagaaaatcgaacctgtgtggaatagtttctacgccccaaacaaactgtag

20 ***Drosophila* Gene Hit** TBLASTN with ORF1: alpha esterase (aE10) gene (U51054)
Human Homologue TBLASTN with ORF1 and BLASTX with U51054: bile salt-
 dependent lipase (S79774)

25 **Annotated *Drosophila* genome genomic segment** AE003671
Annotated *Drosophila* genome Complete gene candidate CG1131 - alpha esterase 10
Human homologue of Complete gene candidate 4e-48 4557239
 ref|NP_000656.1|pACHE|
 acetylcholinesterase (YT
 blood group) precursor
 30 >gi|113037|s

35 **Putative function** alpha esterase
Confirmation by RNAi Only wild type profiles observed

CATEGORY 5: SMALL IMAGINAL DISCS (BLOCK TO PROLIFERATION)

Example 54 (Category 5)

| | | |
|----|-----------------------------------|---|
| | Line ID | 113/20 |
| 5 | Category | 2nd chromosome, small imaginal discs |
| | Reversion | R |
| | Map Position | 50D/E |
| | Rescue ID | EcoR1 |
| 10 | Rescue Sequence 1 | |
| | | CTGAGGCNCTTTGCCAATATGTGTATATTGGGCGGGGNACATGCGTNAATCGG |
| | | TTAAAGCCGCTACTTACATTCTGTTCTTTGCATCTCCCCCATCCACAGCTATAA |
| | | AGCAAGATGAGCTACGCCGCTGATGTGCTGAACCTCGGCCCATTTGGGAGCTCC |
| | | ATGGTGGTGGCGACGCCGAGTTGCGTCGTCCATTCGATCCCACGGNCCATGAT |
| 15 | | TTGGATGCATCCTTCCGCCTTACACGCTTCGCANATCTAAAGGGGGCGCGGCTG |
| | | CAAAGTGCCCGCAAGGAAGTNGCTCCCCCACCT |
| | Rescue ID | BamH1 |
| | Rescue Sequence 2 | |
| 20 | | CCACCTGGTACCACAGCGCTCANACGTGTATGTACACGGATTTTCTGCCGCGT |
| | | GTGTGTAGCGCGGCCCGTGATTGGCTGCAGTCGCGATGGCGGCTAAAACGGG |
| | | CGAAGTCAGTATTTCTCCCTGTGACGANGCGAGCAACGTGAACAATGCCCAC |
| | | TCATTTCAATTGCAAAAATGCCAAAAAGTGCGCGCTTTGAATTGGCCATTTGGT |
| | | TCGTTGCGTTCGTTTGTCTTTTGGTACTTACGTTTGCTTGTGCGATTGTACAAA |
| 25 | | GATAATTGTAGAGTAACGTTAGCAAATTATATTTATTTTGCGCCTGGTTTTTGC |
| | | TTTTCCAACGANCGAGATGTCACAACAGGGTTGTATTANCGTGTGCGGCTGAT |
| | | TCGATATTTGGGATGCCGATTGTCTGAAGCGANGGTTCAACGGGGCTGCCAAC |
| | | TCCCCCGAAAATCTATCNATGGTATTGTGCGCCAAGGGTAAAATAAATAAAAA |
| | | TATGTTAAAACCGCGGAATAAATGGGGGAACCGAAGTGGAAACTGTGGTTCA |
| 30 | | CAGTGCTCTGACTTTCGGGAGCAGTTAATATAGTTGGCATTAAATTCAATTAGA |
| | | GCTCCAAAGTGCTGGTCACAAAGAACGCACAAGAACGGGCCATGAAAAACCT |
| | | GTTGCGCCAGCAGAACGAAAAGTAAAAATTAGAAGAAACCAT |
| | Genomic hit, Accession No. | CSC:AC017131 |
| 35 | Drosophila Gene Hit | rescue sequence: selenophosphate synthetase (ptuf1) (U91994) |
| | Human Homologue | BLASTX with U91994: SELENIDE, WATER DIKINASE 1 (SELENOPHOSPHATE SYNTHETASE 1) (SELENIUM DONOR PROTEIN 1) (P49903) |
| | Drosophila EST | LD46437 (AI514756 similar by BLASTN to U91994 |
| 40 | | selenophosphate synthetase (ptuf1) gene) |

Annotated *Drosophila* genome Complete gene candidate CG8553 selD selenophosphate synthetase

Human homologue of Complete gene candidate 1711372 P49903
SELD_HUMAN
SELENIDE, WATER
DIKINASE
(SELENOPHOSPHATE
SYNTHETASE (1e-159))

5

10

Putative function selenophosphate synthetase

Confirmation by RNAi Only wild type profiles were observed

Example 55 (Category 5)

| | | |
|----|--|---|
| | Line ID | 121/1 |
| 5 | Category | 2nd chromosome, small imaginal discs |
| | Reversion | NR |
| | Map Position | 60B |
| | Rescue ID | BamH1 |
| 10 | Rescue Sequence | |
| | | TCCTGTGCACTCATATTGATTTGCCTTGTCAAGTGGCTAAAGAAATATTAAATG |
| | | TTTGTTATTTCTGTTGCTAGCGCTCCGACAGTCTGGCAGCACTGCTCGCTGTCTG |
| | | ATAGTTCAACTGAGTTGCTGTTTCATCGAACAGAGCTGCCAACTCTATTTTTTGT |
| | | AGCTGGCCAGCCAGGATTGCCAGAGTAAGGCCCTCAAATCAGCTGTTTTTGTGT |
| 15 | | TTTGATTTTATTTTGAAAGTCCTAGTTTAAAATTATGCTTTCTCCGACAGATCA |
| | | GCACAAATAATACTAATAAAGCTCACAATGCTAAGGTTGTGCCTTCCAACCTCG |
| | | AGCTGGATATGTGCGTAAGTAAGGACTTTACGTCTATAAACTTGTTATGTAA |
| | | AGTAAATGTTTGCCTATTGCGCAATTTCTCCAACGAAAAACCCAGAAAACCNA |
| | | AACCCCCTTNAAANTTTGGAATATNCCCAATGAATGCAGCACCCGTGAAATCC |
| 20 | | GTAATGCCTTTGTCCAGCTCTCCAAATTGGTAAGTAAGTCCAAGATCCAAAGG |
| | | AGCCTCCTAAACCCTGCCCTTTCCACAGTACCACCCAGATGTTAAGAGCAATG |
| | | CTGCGTGTCGGAGCGCACAGCCCGATTTGTTTCAGATCTCCGAGGCGTACAAG |
| | | AACCTGATAAAGCCGGAACGGAAGGAAAAA |
| 25 | Genomic hit, Accession No. | CSC:AC020499 |
| | Drosophila Gene Hit | rescue sequence: DnaJ60 gene for dnaJ-like protein (Y11900) |
| | Annotated Drosophila genome genomic segment | AE003463 |
| | Annotated Drosophila genome Complete gene candidate | CG12240 – DnaJ60 |
| 30 | | CG13570 – spaghetti ser/thr phosphatase |
| | Human homologue of Complete gene candidate | CG12240- 4827026 |
| 35 | | ref NP_005138.1 pTID1 tumorous imaginal discs (Drosophila) homolog >gi 3372677 (AF061749) 7e- 08 |
| 40 | | CG1116- 2495728 HYPOTHETICAL PROTEIN KIAA0258(aa) |
| 45 | Putative function | CG12240 : Chaperone involved in protein folding CG13570 : serine/threonine phosphatase |

Confirmation by RNAi

CG12240: Marked reduction in G1 and G2/M peaks
indicating fewer cycling cells
CG13570: Marked increase in G1 peak

Example 56 (Category 5)

Line ID 127/2
Category 2nd chromosome, small imaginal discs
5 **Reversion** NR
Map Position 57F

Rescue ID EcoR1
Rescue Sequence 1

10 GCCGGTGGGCCCACACTTGTNCGCCCGCGCATCGGCTGTCTGTGGGAGTGCGA
NCGAGTCAGATAGTAGATCCGATGCGCTCTCCAGATACTTTTTGAACACTGAA
GAAAACGCGCAGTTGTGGGTGAATTCAGCATCATCAGATTGAATCACACACA
ATCCTAGTCGCCTCACGCGAAGAGAACTATGTCATGATCAGATATCGGTGTAT
GCATTCTATATTATGTACTTCGAAATATGTAATTTATTAAGTTTTTCGCTATACT
15 TTTCAATTCAAATTGGCAAAAACCAATTCAAAGGTTTTCAATATTTTCGAAAAG
CATTTTAGGCTTTCTATGTAACGTATGTTTTTCAAACAAAATATTAGTTTTTGA
AACTTTATTATCGGATAAACAAATGTAAGCCAAATNACAACGTTNTATGATAC
TCCCAAAGATCCGCNCTNTTAAAGTGGCCTAAAAATAGCTGACGCATTAANCC
ATAGGCGCTTCGCTTCTCAAGATAAAACCTGGCGTGCTCAACTCAAGAAACAA
20 ATATGTGGTTATATACATATATACATATATGGGGCATATAACCGATGTGTGAC
GTGACATTGGCTCGTTCTATTCACATACTTAAACACTAAATGCAAACCTATCA
AAAACCNACTACACTAAGCGAAAAACGGCAGANATAGTTAAGGAAAGTGGTC
CA

25 **Rescue ID** BamH1
Rescue Sequence 2

CTTCTTTTCTCAAAAAACGTCGCTCGNGTCCCNCAATCGTTTTACAAACTTCGC
TCGGAACGGACGTGTGCGCGCTCTGAAAGGAAAAAGTGAAAAAGTGTGTGAC
AAAGTGCAAATAAGCCACAACGCGCATGTGAGAAATCAAATTTAATTGAGAA
30 GCATCAAAAATTGTATACATATCGAGCGTATCCACATCGCTGTATGTGTGAGT
GTGCCAGTGCTAGTGTGGTTTTCCCTTTTCGCCGTGGAAAATATGAAAACCTGA
ATGAAAAACTGAATCGCAGTCAGCCAGAGCCGAATTGGAAAAGAGTAACTCG
CATTGGGGACACGAAGAGGTGTCTCGAAAAAGGTAAAATCTTTTACACAGAA
ACGACGCCAGAAAGCGATTAGCGATTTNTGACTATGTGTGAGTGTGTAATTC
35 GGTCTACGGCTGTGTGTCTGCATTTTATTTAACNTTTTGTTTCCCN GTTNGNTC
CACNGTAAAAATAGCTAAAAAAAAGGGCAAGTACTCTTGGCGCGCTCTCCC
TCTCTCTTTGTTGGTCGTGACTGCGACGTACCGTTACGTTAGAAATCGTTTTCA
AGTGGCGTTTCTTTCTTTCTTTTAAATGTGCTGCTTCTTGCTTCTGCCTCTTCTTC
TTGCCTTTGGCTATCTGCTTTGTTTTGAAATACGTCCATGTTATTCCAGTGTCTG
40 TGCCAAATGTGTGCGANATGATCTCTACTT

Genomic hit, Accession No. AC009732

Associated ORF

45 Genscan ORF1 predicted sequence
>/tmp/aaaaafrla|GENSCAN_predicted_peptide_2|456_aa

171

MQTKGPITDADCIRGMACRALAGLARSDRVRQIVSKLPLFASGQLQTLMRDPILQ
 EKRAEHVIFQKYALELLERVSGKTKPLNNPLDPSLSNMHKANVIAQTRIQYNKQQ
 LYQLIFEHLESNGLSQTAAQMLQREVGLPLQTPTRSFHQSPFDYKSLPSGSSSLSRN
 RLRSRMQDVNAAIMGNGDLNRSFGEDSSPAGAGGSNAGDGVSIPIFNSSLNTTQTP
 5 IKIRRTDRSSVSRSIQKQAMEPGGMSVGLAEDGQLHPKRITLNTIVTEYLTNQHSL
 CNNPVTTCPQFDLYEPHKCPDPKPSRLLSSNYNLTSRHARTQAGFNTSRFDRRYV
 HTHFSPWRSIRSADYEDLEFTCCDLAGKYIIVGTQQGDGRVFNMNDGVEQFFSNC
 HNFSVDAIKANRAGDLVITSSFWRTPTSILWSIADDEFKLKLRLPDVTYCEFSQTV
 QDRLLGTQNEVY

10

>/tmp/aaaaafrla|GENSCAN_predicted_CDS_2|1371_bp

atgcagaccaaaggaccattacggatgcggactgtatacgtggaatggcctgtagggccttggcgggacttgctcgtccgac
 gggtcaggcagatcgtcagcaagcttcactcttggcagcggacaactccagacgctgatgcgggatccatactccaggaga
 agcgcgcggaacatgtaatcttcaaaagtacgcattggagttgctagaacgagtgctgggtaagacgaaaccgctaaataatcc
 15 ttggatccatcgctgtccaacatgcacaaggccaatgtaatgccagacacgcacatccagataacaagcagcagctgtatcagc
 ttatcttcgagcacctggaaagcaacgggtctctccagacagcccaaatgctgcaacgggaggtgggtcttccgctacagactcc
 cactacgcgcagtttcatcaatcaccttgcactacaaaagtcttccagtggttagtagctcgtctgtagaaatcgtctgcgaagc
 cgcacatgaagatgtgaacgcagcgataatgggcaatggagacttaaacagaagtttggtagaggactcctcgcggcaggagcc
 ggtgtagcaatgcgggagatggagtcagcataccaaatttagctcccttaacacaacgcagacgcccataaaaataaggagg
 20 acggatagaagttcagttagccgctctatccagaagcaggcaatggagcctggtggcatgctcagttggtcttgcgaagatggtca
 actgcacccaagaggatcacctaaataaccatcgtaacggaatacctaccaaccagcactcgtctgtgcaataatccggtgaca
 acctgcccgcagtttgattgtacgagccgcacaagtgtccagatccgaagcccagccgattgctaagctcgaactacaacctga
 ctatgctggcatgctcgaaccaagccggatttaataaccagtcgctttgaccgctcgtatgtgcacacgcacttttaccatggcgta
 gcattcgatcgccggactacgaggacctagagttcacctgttgcgatttggcgggtaaatacatcattgtgggcacgcagcaggg
 25 cgacggacgagtggtcaacatgaacgatggcgtggagcagttcttctcaactgtcacaactttagcgttgatgctattaaggcta
 agagccggagacttggatcacatctagcttctggcgcacaccaccagcattctatggctctattgcggacgatgagttcaagcta
 aagttgcgactcccgatgtcacgtactgtgagttcagtcaaacgggtgcaggatcgtttgttgggcacccagaatgaggtatactaa

30

corresponds to CG10082

***Drosophila* EST** several including SD04293 (AI532704)

Annotated *Drosophila* genome genomic segment AE003454

35

Annotated *Drosophila* genome Complete gene candidate CG10082 – novel protein with
 homology to enhancer Pi
 uptake

40

Human homologue of Complete gene candidate 1665793 dbj|BAA13393|
 (D87452) Similar to
S.cerevisiae YD9335.03c
 protein (S54640) [Homo
 sapiens] (2e-43)

45

Putative function Putative phosphatase or enhancer of Pi uptake protein

Confirmation by RNAi Reduced G1 and G2/M peaks indicating fewer cycling cells

Example 57 (Category 5)

| | | |
|----|--|--------------------------------------|
| | Line ID | 131/8 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | R |
| | Map Position | 60A |
| | Rescue ID | BamH1 |
| | Rescue Sequence 1 | |
| 10 | CACGATTGCNGGCCCATCGAAGTGTGGGTCTATCGATACTCGTGGGTAAATAA ACAAGTTCTGAACTGCGATTTCTGGGGGTTTGAGGGGTCAATTGTCCCCTGTGT TGGAATGTGTTCTAAATCTACACAAACACTCCCTAAGCTTATCCTAAACTTAT AAATATTGGTTGCTATTTAAACCCCATTTACACGGTTATCCAGCACGCCCCTGA ACTGTGACCCACATCCCCGATTTTAGTGACTAGTTTTATACTTATCGTGGTTGG | |
| 15 | CATTTGGTACACTACACTTTCTTATTCACCTAGATCGCCGACTCCGCGCACGGT CGCGCTCCCGTTCCCGCTCCCGATCTCGGCTGCGACTGCGGTGCGGATCCCGTT CCCGGTGCGGGCGACCGGCGCCTCCANATCCGGATCCCTAANC GGCA NCNGT CNTGGTGGCAATC NNGGAATGTTCCGGGGN NCCNCTACCN CAGTGNAATCAC TGGTACGTCCCACCGCNAACTCCGCCCANTGCGGTTGCCGGAACGGGTGGC | |
| 20 | ANTGCCAATGGGTCGCTGCAGAAGGTACCATCACAGCAATCGCTCACGGANC CCGAAGACTGCCTCTGCCGCCCCGGCTGGGCCACTCATACACGCTACACGGTCG GAAATACTACATTGATCACAATGCGCATACCACGCACTGGAATCATCCGTTGG GAACGC | |
| 25 | Rescue ID | EcoR1 |
| | Rescue Sequence 2 | |
| | AATTGATTTCCGGACATATAAACAGAATCCAGAACTCATCCGGCAGCAGGCTC AGTCAGGCCAGTAAATCCGAAAAGAGAGTAACCAGCAGGAAAAGAGAATCC ACGTAAATACAGAGAAAATGGCTCTACGCGTCCAATTCGAGAACAACGACGA | |
| 30 | CATCGGCGTATTCACTAAACTAACCAACACATACTGCCTGGTGGCCATCGGTG GATCCGAGACCTTCTACAGCGCCTTCGAGGCGGAGCTGGGCGACACCATCCCCG GTGGTGCATGCGAATGTGGGCGGCTGCCGGATCATCGGCCGCCTCACCGTGGG CAACCGCAACGGCCTGCTGGTGCCCAACTCCACCACCGACGAAGAGCTGCAA CACCTGCGTTACANCCTGCCANAACCCCGGAAANATTTATCGTGTGGAAGAAC | |
| 35 | GCCTGTCCGCGCTGGGCAACGTTATCGCCTGCAATGATTATGTGGCCCTGGTG CACCCGGATCTGGACAAGGAGACCGAGGAGATCATCGCGGACGTGCTCAAAG TANANGTCTTCCGCCAGACCATTGCCGACAAC TACTGGTGGGCTCTTACGCC GTGCTGAGCAACCAGGGGGGCATGGTGCATCCCAAGACNAGCATT CAGGAAC AGGACAAC TGTCTCCCTGCTGCAGGTTCC | |
| 40 | | |
| | Genomic hit, Accession No. CSC:AC020517 | |
| | Associated ORF | |
| | Genscan ORF1 predicted sequences >22:13:05 GENSCAN_predicted_peptide_4 357_aa | |
| 45 | MALRVQFENDDIGVFTKLTNTYCLVAIGGSETFYSAFEAELGDTIPVVHANVGG CRIIGRLTVGNRNGLLVPNSTTDEELQHRLNSLPDAVKIYRVEERLSALGNVIACN | |

173

DYVALVHPDLDKETEEIIADV LKVEVFRQTIADNSLVGSYAVLSNQGGMVHPKTS
 IQDQDELSSLLQVPLVAGTVNRGSEVLAAGMVVNDWLSFVGMNTTATEISVIESV
 FKLNQAQPATVTTKLRAALIEDISRSRVAGGGGGGGGGSSSGNSSSSGPSTSRRTT
 RNNAAATAADRPKINEADLEGKSPEEVEMLKTMGFCTFDTTKNRKVEGNDVGEV
 5 HVILKRKYRQYMNRKGGFNRPLDFVA

>22:13:05|GENSCAN_predicted_CDS_4|1074_bp

atggctctacgcgtccaattcgagaacaacgacgacatcggcgtcttactaaactaaccaacacatactgcctgggtggccatcg
 tggatccgagaccttctacagcgccttcgaggcggagctggggcgacaccatcccgggtggtgcatgcgaatgtgggcggctgcc
 10 ggatcatcgccgcctcaccgtgggcaaccgcaacggcctgctggtgcccactccaccaccgacgaggagctgcaacacct
 gcgtaacagcctgccagacgccgtgaagatttatcgtgtggaggagcgcctgtccgcgctgggcaacgttatcgctgcaatgat
 tatgtggcctggtgcacccgcatctggacaaggagaccgaggagatcatcgcgacgtgctcaaagtagaggtcttcgccag
 accattgccgacaactcactggtgggctcttacgccgtgctgagcaaccagggcgcatggtgcatcccaagacgagcattcag
 gaccaggacgaactgtcgtccctgctgcaggttcccctcgtggccggaacagtgaaccggggcagcgaagtactcgccgccg
 15 gcatggctcgtaacgactggctctccttcgtgggcatgaacaccacggccacagagatctccgtgatcgagagcgtcttcaagctt
 aaccaggcacagcccgccacagtgaacgaagctgcgtgcggccctcatcgaggacatatcgcggtcgagggtcgccgga
 ggaggaggaggaggaggcggcggaagcagcggcggaacagcagctccggaccatcgacgtcgcaaggacgacg
 aggaacaatgcggcgccacagctgccgaccggcccaagatcaacgaggcggacctggagggtaaatcgccggaagaggt
 cgagatgctgaagacaatgggattctgcacgttcgacaccaccaagaacaggaaggtcgagggaacgatgtcggaagagtc
 20 atgtaacctcaagcgaaagtaccgccagtagatgaatcgcaagggtggcttcaaccggccgctcgattcgtggcatag

Drosophila Gene Hit rescue sequence and TBLASTN with ORF1: b(2)gcn
 (EUKARYOTIC TRANSLATION INITIATION FACTOR 6
)(X97641)

25 **Human Homologue** BLASTX with X97641: integrin beta 4 binding protein (HUMAN
 EUKARYOTIC TRANSLATION INITIATION FACTOR 6)
 (NP_002203.1)

Drosophila EST GH08760 (AI109537 similar by BLASTN to X97641
 "D.melanogaster b(2)gcn gene.")

30

Annotated Drosophila genome genomic segment AE003462

Annotated Drosophila genome Complete gene candidate CG17611 – bcgn benign
 gonadal neoplasia homology
 to Eif6 translation factor

35

Human homologue of Complete gene candidate 6016331 EUKARYOTIC
 TRANSLATION
 INITIATION FACTOR 6

40

(EIF-6)(aa) and 4504771
 [ref]NP_002203.1[pITGB4BP]
 integrin beta 4 binding
 protein(aa)

45

Putative function eukaryotic translation initiation factor 6 (eif-6)(aa)

Confirmation by RNAi Slightly reduced G1 and increased G2/M indicating block in
 G2/M

Example 58 (Category 5)

| | | |
|----|--|--------------------------------------|
| | Line ID | 135/25 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | NR |
| | Map Position | 24A |
| | Rescue ID | EcoR1 |
| | Rescue Sequence | |
| 10 | ATAACATGGGCNCTGGTTTTTAAGTNAAGCTCTANTNATTGGCCCCCATTCTTA | |
| | NNCTCTCTCGCTCTCTTCTCGCTCTTTCGCCTGCTCTCTCGCCTGATTATTCTGC | |
| | TTGGTCGGCTGATGGTTTTTNGTTTTNATCTGGTGTATTTTCTGCGTAGTTTATG | |
| | ACAAACCGGCTGGTTCTTGTTGTTATTGCCGTATTCTAATATATTTCCCCTATTG | |
| | TTCTTATTTTTGTTGCAGCCTGCACACCTCGGAGGTTCTAGATGATAAGGGGTG | |
| 15 | TAGCGATGGTGGGGGGCTGTCTTGANGGGCTTCTCGCCTTGAGCTCTTGTTTAT | |
| | CTTTGGTCATTTGTTATTGTTTAATGCACGGCAATATTATTGGTAAACAAGTTA | |
| | GCCAACAGCACTAAACGCCAATCGCATTCTTTTCTAAAAACCAAGTCTATTGT | |
| | CGATCTTGCTAGGGAAATGATGATGACTCAGGTGCAATTGGGATCTTATCTAT | |
| | GGCTGTCTGGGAATCAAGAAGTGTTCCCGCAGAATTCGTGAANTACTGCCGCT | |
| 20 | CTCTCCATGGGGCCATTATTTGCACTCGTTTTNCGCGAAATACCATNAATTAGC | |
| | ATAAAGACACGTCGCCGGCAATCGTGACGTAGGCTATNAATGCCTTCTATGCA | |
| | TGTGCNAACTCGCGGAAGCATAGCAATTTGAAGGAACAATATTTTCANTGCAG | |
| | GTTTTAATGGGCTAAAAAA | |
| 25 | Genomic hit, Accession No. CSC:AC014199 | |
| | Associated ORF | |
| | Genscan ORF1 predicted sequences >20:54:54 GENSCAN_predicted_peptide_3 117_aa | |
| | MSASPTARQAITQVMPMITRKVVISDPIQMPEVYSSTPPGGTLYSTTPGGTKLIYER | |
| 30 | AFMKNLRGSPLSQTPPSNVPSCLLRGTPRTPFRKCVVPTELIKQTKSLKIEDQEQQF | |
| | QLDL | |
| | >20:54:54 GENSCAN_predicted_CDS_3 354_bp | |
| | atgtccgcttcacccaccgcccgtcaagccatcacccagggttatgcccatgatcaccaggaaggttgcatctcggatccgatcca | |
| 35 | gatccccgaggtgtactcctcgacgcccggcggaaccctctactccaccactcctggaggcaccaaacttatctacgagcgggc | |
| | tttcatgaagaatctcgtggctccccattgagccaaactccgcccgtccaacgtgccagttgcttgctgaggggaactccgcgta | |
| | ctcccttcgcaagtgcgtgccggtccccacggaactgatcaagcagaccaagtcgctgaagattgaggaccaggaacagtcc | |
| | aactggatctgtag | |
| 40 | Drosophila Gene Hit TBLASTN with ORF1: BcDNA.HL08053 mRNA (AF132557) | |
| | Human Homologue TBLASTN with ORF1 and BLASTX with AF132557: eukaryotic | |
| | translation initiation factor 4E binding protein 2 (EIF4EBP2) | |
| | (L36056) | |
| 45 | Annotated Drosophila genome genomic segment AE003579 | |

Annotated *Drosophila* genome Complete gene candidate CG8846 - phas1 translation initiation factor 4E binding protein 2

Human homologue of Complete gene candidate CG8846 - 4758260
ref[NP_004087.1|pEIF4EBP2|
eukaryotic translation initiation factor 4E binding protein 2 (4e-16)

5

10

Putative function EIF4E translation factor binding protein

15

Confirmation by RNAi Slight reduction in G1 and G2/M indicating fewer cycling cells

Example 59 (Category 5)

| | | |
|----|--|--------------------------------------|
| | Line ID | 141/12 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | R |
| | Map Position | 21A/B |
| | Rescue ID | BamH1 |
| | Rescue Sequence | |
| 10 | GGCTCTTTTCCAAANAGGCAGTTTCTTGNCCTTTCTTGGATTGCTTTGTAGT GAACTNAATCGTTTTTGTGTTGGTTCCTCTGTCGTCAGTCTTGTGAAAATTTTCGTG ATAATAATGCCTGGATAAATANTTAAGCATTGGAACCGGGGGGAAAAAGGG CTAAGTTGTGTGAAGGAAACAATTGAAGTGACCCTTTGTNTATAAACATTCCA CGACGTGTTTCGAAAACAAACAAAGATATGCGGAAACAAAGTGTTAATAAAA | |
| 15 | GAGCNAAAAATAGAGAGAGAGTGTGCGGATAAGCGGTTGAGCGAGATAGAG AAAATTGTTGATTAAAATGTGTGTCNAAATAAAACATCAAGCCGCTTGAACGA ACAGTCAGTTAGTTGCTTCTGATAATAACCATGGGAAGCGGCNCGTGTGCTTC GCTCCTCGTTACTTATAAAATATTTAAACGTTTGCATTCTTCNTATTTCCGAAT TTTTGCNCCCCTGAANCAACTTNGTTAAACTGCAAATAGCAATGCAAACAAAC | |
| 20 | GAATAGAACTGAAATCGACAACNACATGTGAAATTCACAAATCAAATCGCA ATTGTCATCCCAAAGATATAGAACAAGCTATAGGGAAGATANAGAATGTAAG TGCCAAACTAAAATAAACAAACAAGAATAACATTTCCACAGGTGTTTTGCATT TCAAATGCATATTTCCGTGGCGGNTACAAATCTTTTCAAACCG | |
| 25 | Genomic hit, Accession No. CSC:AC017815 | |
| | Associated ORF | |
| | Genscan ORF1 Predicted sequences >17:48:30 GENSCAN_predicted_peptide_2 554_aa MSNKKMFNRTTSVSPGQLHYYHTDFYYSMPLHKTRKMHGVKRVLVFCLMIVIL | |
| 30 | PAILIIMPLHLRKTVPFADVIYPMAESDIIIRAGISSIFCSKHTLRMNSNFNAFQLRNK PEIATNRKHRLKKSMTLPDDTLEYWGFLLKGAKVRVKFCSRYDGSRLIIHGHR ELNLCGLTDHNKNKLGANAYAKGHEQVQVFFEDNVEITEEKGNQDVLMEHENHG GEDLTEDIPQPQVNIPVKQNNISIQPKLIRKKLKKGTIHHGEHDMHAITDLQGSHHT EHILNHHDHSSNSPAHHNSTAHHREHSSNITNEETSRNHIRNEDEDPDQNSSKTH | |
| 35 | YSAESPPIRERLKRHNRAHRNQKRQDLYDTLYKRSKRENVYDRKTIHGGNAIN FTETDESNSVSSFETGLFQCFNGMILLQEFFRPKNECSNPHIMDTSPNKSSMVVHN VIEDGYYYYIFYSDNDHVQNEIHAIFDIYKPTYQYSNMSESQSCLNTTNCTFNISFL SDEIVVVEVPTRDIEHEEDDITNLISTCHPRSEIYAIFPITVLVLILCCSFL | |
| 40 | >17:48:30 GENSCAN_predicted_CDS_2 1665_bp atgtccaacaaaaagatgtcaacaggactacgtcagtaagtcctggacagttgcattattatcacacggatttctattactcaatgcc ggatttgataaaacccgcaaaatgcacggcggtgaaaaggggtgctggtttctgcctgatgattgtgatactgccggccattcttctc attatgccgctgcatttgcgaaagacgggtgttggcgacgtcatctatcccatggcggagtcctgatattcattgagattcgggcagga atctcgtcgtatctttgctcgaacacacactgcgtatgaactccaatttcaacgcttttcaactacgtataaagccggaaattgcgac | |
| 45 | gaatcgcaagcacattaggtgaagaagtcgatgacattgccggatgatacgttgaatactggggcttcttcttgcgaaagggtgc caagggtgcgagtgaattctgctcccgctacgatggatcccgcatcctgatcatccatgggtcacaggagcttaattcttgcggtct | |

gaccgatcacaataagaataagttgggcgccaattatgccaaaggtcacgaacaggtgcaggtgttcttcgaagacaatgtggag
atcacggaagagaagggcaaccaggatgtgctaattggagcacgagaaccacggcggagaggatttgactgaggatattccaca
gccgcaggtgaacatacctgtcaagcaaaacaattctatacagcctaagttaattaggaaaaaactgaaaaagggcacaattcatc
atggcgaacatgatatgcatgctataacagatttgcaaggatcacaccatacgggaacacatattgaatcacatgatcacagctcta
5 attctccagcacatcatcacaatagtactgcccatcatcgggagcacagttcgaatatcacaacgaagaaactagtcgtaatcaca
tacgaaatgaagatgaagatccagatcagaattcaagtaagaccattatagtgcggaaagtccgcctcacggggaacgtctcaa
aagacacaatagggtagcccataggaatcagaagagacaggatctttacgatacgtttataaaagatcaaagaggggagaatgtc
tacgatagaaagacgatccatggaggaaatgctataaattttacggaaacggacgagtcgaattcgggtgtccagctttgagacagg
actatttcagtgtttcaatggaatgatcctgctgcaggagttcttcaggccaaaaaatgaatgctcaaatccgcacataatggacactt
10 cgccaacaagagttccatggtggtgcacaacgtcatcgaggatgggtactactattatataattctacagcgacaatgatcacgttc
aaaacgagatccacgccatattcgataattacaagccgacgtatcagtactcaaacatgagcgagtcacaaagctgtctgaatacc
acaaattgcacattcaacatcagtttcttcggatgagattgtggtggtggaggtccaacacgggatggtatcgagcacgagga
ggacgatataaccaatctgatctccacctgtcatccgcgcagcgagatatacgccatctttccattacgggtgctggtgctgatcctt
15 gctgctccttctgtag

corresponds to CG9524

| | | |
|----|---|--|
| | Annotated <i>Drosophila</i> genome genomic segment | AE003623 |
| 20 | Annotated <i>Drosophila</i> genome Complete gene candidate | CG9524 - novel His-rich |
| | | protein |
| | Human homologue of Complete gene candidate | none |
| | Putative function | No homologies which indicate function |
| 25 | Confirmation by RNAi | Reduced G1 and G2/M peaks indicating fewer cycling cells |

Example 60 (Category 5)

| | | |
|----|--|--|
| | Line ID | 146/2 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | NR |
| | Map Position | 26B |
| | Rescue ID | EcoR1 |
| | Rescue Sequence | |
| 10 | | TTTNATCCAAACTGAGANACTNTTGGCCCCAAAACTGAAAACCTCGGACTCGGG CGCGTAAGGGAGTCGGTCNTCGGGAGTCGGTCGTCTTTTGTTGATCTTGAGAC TGAAATTCCAATTGTTGATTTATCTCTCGGCTGCTGCGCCGCGGCTGCGCTGCT GCAGCGCAGTCCCACTCCGATTTGACCAGCGACCAAGTTTATAAACTTTGAG CCAAAATGCAGCGGCGCACAGTTGTTACCAAAACGTTGCACGCGTCGTGGCCC |
| 15 | | TCATCAAAACAAAAAAAAAATATAAGCGAAAATGAAAACGAAATTCGGTTA ACGTCAACAGAAGCTGACAAAAGGCAGAAAAGACCGAAACAAGTTGCAGGG CCAGAGTAAGCCAAGTTAAATGCGAAAGAGAAGCAAGAGNCAAGAAGAAAN AATGGGCACTACATACATATATTATAGCCAGCTAATCTGTTGTGCAGTGC GTT TTATCAGCCNNCGAAAAGAAAACGAAAACGAAAAGTCGGTCCAAGTTCGGAC |
| 20 | | TCAAAATCCAAACAGAAGAGACTCCATNCCATCAGAGACACGCGGATCTCAT CTCGGTAATGTCTCAATAAAAGTAATCTTAACTGCCGCCGGGAATGTTGGAAA AAGTGAAAATTGAAGCGCTTAACGTGTTTCGAAATACGATACATGAGAAGTCC CAAAAAAAAAAAAA |
| 25 | Genomic hit, Accession No. | CSC:AC019865 |
| | Drosophila EST | GH19286 (AI388389) |
| | Annotated Drosophila genome genomic segment | AE003481 |
| 30 | Annotated Drosophila genome Complete gene candidate | CG11353 - novel with weak homology to sugar acetylase? CG7525 - tie receptor protein tyrosine kinase. |
| 35 | Human homologue of Complete gene candidate | CG7525- 4e-23 4557869 ref NP_000450.1 pTEK TEK tyrosine kinase, endothelial >gi 464868 sp Q02763 TIE2_ |
| 40 | Putative function | Sugar acetylase and receptor tyrosine kinase |
| | Confirmation by RNAi | Both gave a reduction in G1 and increase in G2/M peaks indicating arrest in G2/M |

Example 61 (Category 5)

| | | |
|----|---|--------------------------------------|
| | Line ID | 155/13 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | R |
| | Map Position | 21B |
| | Rescue ID | BamH1 |
| | Rescue Sequence 1 | |
| 10 | GNTTTAGTCCNCTTTTGANAGGGNCTTGGNGNCTTAAANAANNAAAAAAGGG | |
| | GNCCCGGCNCCCAGCAAANAGNNTAAACTTGAATGGTTTAATTCGAAAATC | |
| | TTTTAGAAATGTCGCCTAATACCTTATCGGTATAGAGTTCACCTCGTCTCCTAA | |
| | TCCATATTTTAAGATATCAATATCTATTAACAATTTTATCGTATGATTAGAAA | |
| | TTCGCATTGTTTTATTATTTTCGACCTTTGGGCTTTACATCGACAGCTACTCTCTA | |
| 15 | TCCAGACAGGAGACTGGGAGAGAGAGCACGATGCTGTCTGAAAGCATGAATG | |
| | ATGGATGCTGTGCCTATGTGCGATATGCACGTTGCCTGAGCTAAAACGAAACG | |
| | AGATTATTAATCTATCCGCAAGATTCAGATGCTGATTCCACATGAGTGAGCGA | |
| | GTCCGTGAGTGGATATTGCTCTCTCCGAAATGCATGCATGAGTGAGCAGGGGG | |
| | GCTTCAATCGCNCTNTCGATNTGCGACAGNGACATNTTTTTATCTTCGACNAT | |
| 20 | GCNCTCNCTCCCTCCCACAGAAATCTTGCGCTNGNTCTCCGANNTNGGGNTNG | |
| | ANGGCNCTCTTCTCTNTCCTTAAATTGGGANTTNNCTTTTTTCNAANAAGGGN | |
| | NAGA | |
| | Rescue ID | EcoR1 |
| 25 | Rescue Sequence 2 | |
| | AATCNTTTTNTCCATTNGGCGNCTTNCTCAAACATATTCACATTTGGNCCCAA | |
| | CGGCGTANGACTTNATCTCACGATTGTTTGGTTTCCTACTCTCCCGCGCTCCCT | |
| | CTCTTCTGAGTCTCTTTCTGGCTGATTCGCATTCGATTTTAGCCGCTGCCATCG | |
| | CCGTTGTTTTGCCTACCTATGTGTGTGTGTGAGGAGTGTGTCTTGTATTTCA | |
| 30 | CGCAATGCGCTCCGCTCATTATTTGTTTGANCGCCGCGGTGTAAAGTTGTAA | |
| | AAAGTCCAAGTGCTCGTGGAACCTCGATGCAAGACGGGGAAAACGAAACGCG | |
| | ATAAATCGTGAGAAAAGAGAGTGCGCTAAAGGAAGAGGGAGTGATAATCAN | |
| | ACGAAATGGAATAATGTNTTTGCAGAGGCNACAACAATGCAAATAGTTG | |
| | TCATTGAGGCGCAATGAATGATAATTAGTGCTTANTTGAAATCATAATCNTGA | |
| 35 | AGAAAGCGTAAAGCTCGATTNTGGCAATNTATTCTTGATTACCANTGAGTCTG | |
| | TGATATTGCCGTGTGTNCCGAAAATGGANGTTATNAAACCCATGGACTTCAGC | |
| | ACCTTCTCCGCGTTCTGCGAACATCTTAACAAATCTCCACAAAATTGCAGCAA | |
| | CAACTGCANCGACGGTACCGCCAATAANCAATGGAAAANGCATTATTTG | |
| | GAGGTAANAGCNAAAAATACCAATNTTCCAATGCGAAATTGCNAGCNTGG | |
| 40 | | |
| | Genomic hit, Accession No. AC004274 | |
| | Annotated <i>Drosophila</i> genome genomic segment AE003590 | |
| | Annotated <i>Drosophila</i> genome Complete gene candidate CG13693 - novel | |
| 45 | | |
| | Human homologue of Complete gene candidate 6e-05 4507659 translocated | |

promoter region (to activated
MET oncogene)
>gi|1730009|sp|P12270|TPR_
HUMAN POOR MATCH

5

Putative function No homologies to indicate function

Confirmation by RNAi Only wild type profiles observed

Example 62 (Category 5)

Line ID 162/24
Category 2nd chromosome, small imaginal discs
5 **Reversion** R
Map Position 55C

Rescue ID EcoR1
Rescue Sequence 1

10 TTTTNTTTCANGGNTCTTTGCNCATAAAAANACACGNGCCCTCNTGTCCATTTCAC
ATTTTACTTGGAGTCGGTAACGTTGAGTTCCGCGTCCGTGCGTTCTGCCTTCCA
ATACAAAGTCTGGTGTGAATCTACCAAGCATTCCAGTGNGAAAATCAACTCAC
ATTGCTCGGTGATCCNTGCGGCGGTATNATCGCACCCGGAATTGCATAAGTTG
CGGNGAGCGGAAAGAGAGTGCACGGATTTCNCNGTTATCNAAGGGCCGGCANC
15 NGTGGGGCGGCGACGGNAGAGCACGCAGAANAANAATANANTGNNGTGGCG
AATTNAAAAATANNATNAAAGAAAATTTCGGGCGCTAATTTTTCTTCAAATTT
GTGTGCGGTTCGGCGAAAAACAACGTGTTTTTCNATGGTTGATAATACACACGG
ACGGNNCACTCGCGCTCACCCACATAGTCACNAAAGTCGGCGACGTCGACGA
CCCNACACNCTCACATANGGACNTTTAATCCCGTNCATNCGTGTAGCGTNCNTA
20 TTTAACCNTNTCTGTCCATCGGAACGCNCGCNTTCTCGCCTTCNTTCTNCTTTA
CTTTAATTTCTTATTTNNAAGGGGNAGNCCNATCTTTTTNCCTNTCNNTGCCNT
TTAANNTCATCCACANCCTCNCTTTNTCNTTCTCCTCCNCCTTNTNTTCTTTTCNTC
TTNCTTNTGNCCTTGCCTCGTTCTTTCTCTTCNTCTCCTTNCCCTTCTCCTCCTTT
TTTCTCCTTCCCCC

25 **Rescue ID** BamH1
Rescue Sequence 2

AAGNCNCCTTGGCCGNNTTNAACGGNAANTAANCCGGGNCCNCGGGNCNCGA
TAATCAGGTCNANCCTTGTGCCTACCACCACCAAATTGAAAAAGAGCNAAGA
30 TTCTCTAAGGCAAAAAAATCCCAATCTGTGGAATTTCCGGAAGCGAGAGCAC
ATTCAAAGCTACCAGTTATCAGCGAGCAGCATGTCTAAGCTCAGGAACCTGTT
GCCACAATCTTTGGCGGGAAGGAGGCACAGAATCCGACACCCGTCGAGGGA
CGCCTGGAATAGGACGCAGCTCCCGTGGACGACAACGAACCNGATTACTACT
ACTGCGGAGCCATGGCGCTGCCCTCCACCGCTGGCACGCCACAGCCTCCTCG
35 GATCTGACCGAATCCGTGCTGCGCGAGCTCAGCGACCCAACTACAATTCAAT
GGATGTGGTGCTTTCNCCCTNTTTTCCGGGCACTCTCAGTAACGTCCAGACAA
ACAACACCATGAACGTTACNGCGCCAGCAACAGGTGGTCATGAACTTCTCG
AATGCCAATAATCTGCACTTCGGCTCCGTCTTCAACTTCAACCAAACTTGAG
CGCCTGCNGCTCNCGAANGGGTTTACCNGTTCGCANAAGAATCGGTTCGCCTC
40 TCCANACNGT

Genomic hit, Accession No. CSC:AC012981

Associated ORF

45 Genscan ORFs: ORF2 predicted sequences
>18:26:17|GENSCAN_predicted_peptide_7|1320_aa

MEETNNATTIEQQPIALINGQEQVANEQQPSSPTSVATPTSTTSGGTGNATPAFSY
 DDLFPALPANTSASQSGASGSTLARVTSSQKTHIVHVPCKERKSTESEKFGESES
 KRICQQITKETGAQIEIASRQVTVPREHFRVILGKGGQRLREIERVTATRINIPSQSD
 ESEFITIAGTKEGLAQAEQEIQLSAEQYKKSSDRITVPKVYHPFIVGPYSENLNKLO
 5 EETGARINVPPQQVQKDEIVISGEKDAVAAAKAKVEAIYKDMKKCSTVSVEVAK
 PKHRYVIGPKGSTIAEILQLTGVSSEMPPNDSPSETITLRGPQVALGNALTVVYQK
 SNSVKSVEINAAHWIHKYVFGRKGANMKQLEEDCPNVNVCLEDKIKLEGDPEN
 VDRAVAYLSEIKNYEENFTFEVMTVNPSSYKHIIGKAGANVNRLKDELKVNINIE
 EREGQNNIRIEGPKEGVRQAQLELQEKIDKLENEKSKDVIIDRRLHRSIIGAKGEKI
 10 REVKDRYRQVTITPTQENTDIVKLRGPKEDVDKCHKDLLKLVKEIQESSHIEVPI
 FKQFHKFVIGKGGANIKKIRDETQTKIDLPAEGDTNEVIVITGKKENVLEAKERIQK
 IQNELSDIVTEEVQIPPKYYNSIIGTGGKLISSIMEECGGVSIKFPNSDSKSDKVTIRG
 PKDDVEKAKVQLLELANERQLASFTA EVRAKQQHHKFLIGKNGASIRKIRDATGA
 RIIFPSNEDTDKEVITIIGKEESVKKAREQLEAIKECDEVTEGEVSVDPKHHKHFVA
 15 KRGFILHRISEECGGVMISFPRVGINSDKVTIKGAKDCIEAARQRIEEIVADLEAQT
 IEVVIPQRHHRTIMGARGFKVQQVTFEFDVQIKFPDRDATEPVEGLTNGGSGENG
 GENEGQEGEQEVEKEAEQEPVRQCDVIRITGRIEKCEAAKQALLDLPIEEELSVPF
 DLHRTIIGPRGANVRQFMSKHDVHVELPPSELKSDVIKVCCTPARVAEAREALVK
 MIEDYEADRADRELRSFVLQVDVDTEFHSKLIGRHGAVINKLRADHDVIISLPKRD
 20 EPNDRIISITGYQANAEAARDAILEIVGDPETLHREVIEIDKRIHPHLIGQRRRTIRKII
 EDNKVNIKFSADDDNPNSIFISGKIEDVENVKELLFGMAEDYERDYLDNVAIAPPTI
 GAFLTGFWIRCRRRCQRRERIRHQRRTVGEAKAGQKPDCAQHSVAGGLPALRCWRG
 SGGLHAYHLRVGPQKLSASGRVSRSPAVAAILQVGVRRGSELEMDQEQKLELE
 LELDYRAMSGRAAAVVRTSL

25 >18:26:17|GENSCAN_predicted_CDS_7|3963_bp
 atggaggaaactaacaacgcaactaccatcgagcagcagccatcgctctcattaatggccaagagcaggtggccaacgagca
 gcaaccatcctcgccaacttcagtggccacgcccactagtagcggcggaactggcaatgccacacccgccttagctac
 gacgacctgttccggccctgccggccaacacttcggctcaatcgcaatccggagcttccggttcgactctagctcgtgtgacgag
 30 ttcccaaaaaactcatattgtgcatgttccctgcaaggagcgcaagtccacggagtcggagaagtgtggcgaaggcgagtcgaag
 cgtatttgcagcagatcaccaaggagaccggagcccagatcgagattgccagtcggcaggtgaccgttcctcgggagcacttc
 cgcgtcactcctcggaagggtggccaacggctgcgcgaaatcgagcgtgttactgcgacgcgcacatcaacatcccagccagag
 cgatgagagcgagttatcacgattgccggaaccaaggagggtattgccaggccgagcaggagatccgtcagctgtcagccg
 agcagtacaagaagtcacgaccgcacacggtgcccgaagtattaccatcccttcacgtggggcccctacagcgagaacctaaa
 35 taagctgcaggaggagaccggcgctaggatcaacgtgccgcccagcaggttcagaaggacgagatcgtcatctcgggagag
 aaggacgcggtcgcagcggcgaaggccaaggtggaggccatttacaaggatatggaaaagaagtgtctaccgtcagtggtga
 gtagctaagcccaagcaccgatattgattgtccgaagggtccaccatcgccgagattctgcagttgaccggtgtgtctgtag
 agatgcctcccaatgactccccctcgagacgatcaatttgcgtgggcgcgaagtggctttgggaaatgccttaaccgttgtctac
 caaaagtccaactcggtaagtctgtggagatcaatcgggcacattggatccacaagtattgttgcgtcgaagggggccaaca
 40 tgaagcagctggaggaggactgccccaacgtgaacgtgaattgcctggaagacaagatcaagctggaggagatcccagaaa
 cggtgacagggtgtgacctactgtccgaaatcatcaaaaactacgaggagaacttcacattcgaggtgatgacgggttaactcttc
 gtactacaagcacatcatcggttaaggctggagccaacgtaaatgcctgaaggatgaactgaagggttaacattaacatcgaagag
 cgcgaggggccagaacaacatccgtatcgagggtcccaaggaggagtagcggcaggcgcagcttgaattacaagaaaaaatcg
 aaaaactggaaaacgaaaaatcgaaggatgtgatcatcgaccgcgtctccatcggttctattatcgagctaaggggcgagaagatt
 45 cgcgaggtgaaggaccgtaccgccaggttacaatcacgatacctacgccccaggagaataccgatattgtgaagctgcgcgg
 acccaaggaggatgtggacaagtgtcacaaggatctgcttaagctggtcaaggagattcaggaatcgtcgacattatcgaggtg
 cccatctttaagcagttccacaagttcgttattggcaaggcggcgctaacaatcaaaaagatccgcgatgagaccagactaaaat
 tgatctgcctgccgagggtgacaccaacgaagtgtatcgaatcaccggcaagaaggagaacgtgctcgaggcgaagggaacgta

tccaaaagattcaaaacgagctttccgacattgtcaccgaggaggtgcaaatcccgcccaagtactacaactcaatcatcggcact
ggcggcaaaactcatctcctcgatcatggaggaatgcggtggtgtttctatcaagttccccaacagcgactccaagagcgataaggt
cactattcgcggtcccaaggacgatgtggagaaggctaagggttcagctattggagctggccaacgaacggcagctggcttcctt
accgccgaggtgcgcgcccaagcagcaacaccacaagttcctgatcggcaagaatggcgcttctatccgtaagattcgcgatgcc
5 actggtgcccgcattatcttcccttcaaacgaggacactgacaaggaagtatcaccatcattggcaaggaagaaagcgtaaaga
aggcccgtgagcagctggaggcgatcatcaaggagtgcgacgaagtaaccgaaggtgaggtttctgctgatcccaagcaccac
aagcacttcgtggccaagcgtggcttcacctgcaccgcatttcggaggagtgcggcggtgatgatctccttccccgtgtcgg
catcaactccgataaggtgacgatcaagggtgccaaaggactgcattgaagcgccccgccagcgcatcaggagatcgtcgccg
atctggaagcgcagaccaccatcgagggtggtgattccacagcgtcatcgcaccatcatgggcgccacgtggatttaaggttca
10 acaagtcacctttgagttcgtatgtgcagatcaagttccctgatcgtgatgccaccgaacccgtcgagggtctgaccaacggagge
agcgggagagaatggaggcgagaatgaaggccaggaggagagcaggaagtagagaaggaagccgaacaggagccggttc
gtcagtgcgatgttatccgaatcacggggcagaattgagaagtgcgaggccgccaacaggctctgcttgatcttatccccatcgag
gaggagttgtcggtgcctttcgacctcatcgtaccatcatcgaccgcgcggtgccaatgtgcgtcagtttatgtccaagcacgat
gtgcacgtagagctgccacctagtgccttaagtcggatgtgatcaagggtctgcggtacgcccgtcgcgtcgccgaggcccgcc
15 gaagcgctggtgaaaatgattgaggattacgaggctgatagggccgatcgtgagctgcgctcctttgttctccaggtggacgtaga
tacggaattccattcgaagctcattggtcgtcatggcgctgtgattaacaagctgcgtgccgatcacgacgtcatcatttcgtgcct
aagcgggatgaaccaatgaccgcacatctctatcaccggctaccaggccaatgcggaggcagcccgcgatgccatcctaga
gattgttgccgacccccgagacacttcacgcgaggttatcgagatcgataaacgcaccccccacctcattggccaacgccga
cgcaccattcgcaagatcatcgaggataataagggtgaacatcaagttctcagctgatgatgacaaccccaattcgatcttcacgt
20 ggcaagatagaggacgttgagaacgtcaaggagttgctcttcggcatggctgaggactacgagcgtgactacttgataacgtg
gcgatagcgcgcccaacgattggtgccttcctaactgggttctggatccgatgccgcaggtgccagcgagaacggattcgtcatc
aaagacgcaccgtgggagaagcaaaagcaggccaaaaacctgactgcgcccacactcagtcgcaggaggacttcccgcact
tcgtctgtggcggggctccggtggcctccacgcctatcacctccgtgtggggcccccaaaaactaagtgcacggggccgagtgtc
ccgatcgccagcagtagcagcaataactacaagtcgggggtgcgccggggatcgagagctggagatggaccaggagctggagca
25 gaagctggaactggaacttgaattggattatcgggcaatgagcggcagagcagcggcagtcgtgcggacatctcttag

Drosophila Gene Hit BLASTN with rescue sequence 1: dodeca-satellite protein 1 (DDP-
1) (AJ238847). TBLASTN with ORF2:dodeca-satellite protein 1
(DDP-1) (AJ238847).
30 **Drosophila EST** GH20785 (AI389573), LP07358 (AI294065)

Annotated Drosophila genome genomic segment AE003799

Annotated Drosophila genome Complete gene candidate CG5170 - Dpi dodecasattelite
35 DNA binding protein
CG5576 - Bg5 involved in
cytoskeleton organization and
biogenesis which is putatively
a component of the plasma
40 membrane

Human homologue of Complete gene candidate CG5170- 4885409
ref|NP_005327.1|pHDLBP|
high density lipoprotein
45 binding protein
>gi|2498434|sp|Q00341|HB

5

CG5576- 2e-07 4506539
ref[NP_003795.1|pRIP|
UNKNOWN >gi|3426027
(U50062) RIP protein kinase
[Homo sapiens]

10

Putative function

CG5170: DNA binding protein (homology with Scp160p, a new yeast protein associated with the nuclear membrane and the endoplasmic reticulum, is necessary for maintenance of exact ploidy)
CG5576: death domain containing protein, possibly involved in signal transduction

15

20

Confirmation by RNAi

CG5170: Reduced G1 and G2/M peaks indicating fewer cycling cells and more polyploidy
CG5576: Loss of G1 peak

Example 63 (Category 5)

Line ID 40/2
Category 2nd chromosome, small imaginal discs
Reversion NR
Map Position 39B

Rescue ID BamH1

Rescue Sequence 1

TTTTTGCCTCCGCTTTTAAATTAATAAAAAAATGTNTGTTTNGCCCTGGAGCTCTCG
 10 GTCTGTTAGCGAGCGTTGCCACCTTTCTGCGAGCTGTTGCTGCACACTGCCACT
 TTACGAACACAGCTCTGATAGCGGGACAAAATACGTCAAGGCAGCGACGGTG
 GGTTACTAGTGAATTTGGAACGGTGGTCTTAAGACGTACTGGTCTTTTATATTT
 TCATTATTTTTTAAATTGTCGCTCATTTACCAATAAACCTTTTTACTTTTTCTCTG
 ATAGTCCGAAGTCAGATCAAATAGGAAGTTTCACAAAAAATTTTCATCCAGAG
 15 AAAATACGCCGACGCTATTCGAGTTTTTTGTATTTCGTTAACCGGGAAAGAATA
 GTTCGAATTCGTTTCGCACTTTATCGATAGTAGATTGCTATTATGGAGCCCACTA
 GTAAATTAATTAAATTCCAGACTGATAAAAGCGATCAACTTTTGTTAATGGGT
 TTAANTCTATAATAATNCTTAGTCCAAATTGTNTCAAAGTAGTCGATAATTTAT
 AATAACAGTTTTAGATGACCTCTAGGAAATAACTAATTACCCACATNCTTCAA
 20 GAAAGTGTTNCAATTTGTNCTATAATTAAATAACAGTTGTATTAATTATGTTG
 TNATTGTNACTCATAATACAAATTAACAATATAAACACACATAAATAAGAG
 AATTGGAATATTTTGTCTCAGATTAGATTTNCCAC

Rescue ID EcoR1

Rescue Sequence 2

AACGGGGGGGCTTCCGCGNCNCCAAAACGCAATNTACCGTTCATGCTGTGAAG
 CGAAAAAGAGTGGTAGCGCCTACCNTGGCATATGTAGTTAAATCCGTGAAAT
 AAGTGAATAAGAATATATGTATGTACTTAATTCGAAAACCTTTTCGCCGTCAG
 CACAACGGGTGAACGAGAGAGCGGAAGTGGAGTTTTTTGTGGCGGGTCGTCT
 30 CGCTCGCACCGCAAANGTCGTCCGTGGCTGCGTGTATGGGTGTGTGGAAAAA
 GCGTCGAGGTGAATGTGGATTTCTAACCACACCAGCATTGCAAAGACATTGAT
 TGATATTTAAAGCTGCAGCAGCGAACAAAGCAAATCCTAATTTTCGGCAAAGTT
 TAAGAATAACGAGTGACTGGGGCGCGCGCAATAAGATAAAATTGAAGGTTAT
 CTGTGTGCGTGTGAGTGACCGTNTACCAGTGTGTGTGTGCGANCGTCCATTGT
 35 AAACAAAAACAAGTGTGTGAGCGGAGAGAAGAAAGGGAAAGAGAGAAAG
 AGCGAACAGACTGGCGAGAGAAAAAAGAGATGCCACAAANAAAGCAGCGCA
 CAAAGGAAAGCTGAAAATTTTANTAAATCTGCAAAAGTGAAGAAAACCAAA
 GAACCCGCAGTCNTGTTAAATAAAACCCAGANTCCAAGAAACNTTAAAAGAA
 GCAGTGCAAACAAACTGGTGCTNTGAATGCGGTTTATTTTGAAAAAAAATGCA
 40 ATTCGGTCCGATGGAA

Genomic hit, Accession No. CSC:AC014744

Drosophila EST several including LD46342 (AI544109 BLASTN similar to mRNA
 45 L07550)

Annotated *Drosophila* genome genomic segment AE003669
Annotated *Drosophila* genome Complete gene candidate CG8678 - novel with ankyrin
homology

5
Human homologue of Complete gene candidate CG8678 -gi7661580
B69CEC399B56F35C
|ref|NP_056425.1|DKFZP434J
10 154 protein [Homo sapiens]
(2.20E-85)

Putative function Novel protein with ankyrin domains, unknown function

15 **Confirmation by RNAi** Reduced G1 and G2/M indicating fewer cycling cells

Example 64 (Category 5)

| | | |
|----|---|--------------------------------------|
| | Line ID | 55/12 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | NR |
| | Map Position | 49C |
| | Rescue ID | BamH1 |
| | Rescue Sequence | |
| 10 | TCTCATGNTCAGGGGGCCTTTACNATGTCAAAGAGCAAATTGTCCACAGGGCA | |
| | GCAACCGCAAGTGAGAGACGGGTGGAAAAGTGGGCGGCATGACCATGAATGA | |
| | AAGCCGCGACCGGCAAACGTGGCCCGCCACAAAGCGAGCATTTTCACATTTT | |
| | AACTGTCTGGACATTTTGTAAAGTTACACCAAGGCAATGATACCAGTAAAAAAG | |
| | AAGAAACAATCATTTTTGAATAGATTAATCACCTGATTAATGTTGGTTGTATGT | |
| 15 | TGATTGTAGGTGTTTTAATATACAATGTCTCTATTACTGCTTTCCTTTATTCAA | |
| | AGCCATGTGTAAGTGTAAGTTCTCGATTTTCGGCTAGATTTTGAAGTTCTGCCAT | |
| | TATCAATTAAAAGTCCAGTTCCTCTATAAATTGGTAATAAAATAGCTCTTTACA | |
| | GCCAAGTATATGTGCAATTTTGTAAAGATTAAANGTCCAAATGTTGTGAACCTT | |
| | TCCTGGCCCTGAATTTTAAAAAACCATTAAATTGGTCCCATTTGACATTAAATG | |
| 20 | TTCTATGTACATTAATATGACTTTTTGTGGATGGTTTTATAAACAAGCATTACT | |
| | ATATTCTAAAAATCAAGGATAAAGGACNAGCTTTACAGGAGGTAACATTCTTA | |
| | TTGTACGGCTTTATTTTCTTATACCCATAAGAGCATACCACTAGGATCCGTCGA | |
| | CCTGCAGATCTCTAAAAACTTGCCTTTGCTGGCGTTTTCCATAA | |
| 25 | Genomic hit, Accession No. AC007085 | |
| | Associated ORF | |
| | Genscan ORF1 predicted sequences >21:54:11 GENSCAN_predicted_peptide_3 108_aa | |
| | MGLVTAAFKLKRKDIQDRYQHDINRICHTRSTAHTAYAHFAEHLRRSPRQRFVN | |
| 30 | GKGAALVLILLVSAARQFSGSTGAYKLGNRVGKVEGEQQEYKLQDRTTTHFCGN | |
| | >21:54:11 GENSCAN_predicted_CDS_3 327_bp | |
| | atggggctggttaaccgcccgccttcaagctgaagcgcaaggatatccaggacagatatcagcatgatattaaccgcatctgccaca | |
| | cacgtagcacggcacacacggcgatgctcattttgcggagcatctgttgcgacgaagtccacgtcaacggttgtcaacggcaa | |
| 35 | aggtgctgcgcttgctcatcctcctcgttctgcggctcgacaattttctggctcgacaggtgcctacaaactgggtaatagagttg | |
| | gaaaagtagaaggggaacagcaggaatacaaaactacaagacagaacaacacattttgtggcaattaa | |
| | Corresponds to CG8732 | |
| 40 | Annotated <i>Drosophila</i> genome genomic segment | AE003836 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG8732 - l(2)44Dea |
| | | homology to fatty-acid- |
| | | Coenzyme A ligase, long- |
| | | chain previously described |
| 45 | | spindle/chromosome |

| | | |
|----|---|--|
| | | abnormalities in neuroblast squashes |
| 5 | Human homologue of Complete gene candidate | 1e-171 4758330 ref[NP_004448.1 pFACL3 fatty-acid-Coenzyme A ligase, long-chain 3 >gi 4165018 dbj BAA371 and LCFD_HUMAN LONG- CHAIN-FATTY-ACID--COA LIGASE 4 1e-157 |
| 10 | | |
| | Putative function | Fatty acid CoA ligase |
| 15 | Confirmation by RNAi | Only wild type profiles observed |

Example 65 (Category 5)

Line ID 6/7
Category 2nd chromosome, small imaginal discs
Reversion NR
Map Position 28E

Rescue ID BamH1
Rescue Sequence 1

10 TATNAATAATCATAGGGCTCTTGCTCTTACGTGTAAGGCCTGCCCCCTCTNCCA
GTCTATATACAAAGAAAAACACACACACACTGGCACACTGGTGTTCGCATATG
CCAAAGCCGAGTTAATTTCACTTTGTTTAATCTATCGTTTGGTGTTCATTTGCATTT
TTTAACCGCGCAAACGGTATTTGCGCGTTCCTTACTTTGCGATTAT
TGCACCGCTTGGCTGTGTTTTCGCAATTTCTATCTTGATTTTCATTGGTATTCACG
15 CGTAATGTAATTCTTAGCAGCGTGACCGCGCCGATAACGATAAAAAATACCAC
GGGACCAAAAAATAAATACCATATGATACCACTTCAGGGAAAAGAAATCCTAT
TTAATACCACTCACTTTAAAAATAAGTTTTTAAAAATATATATNTTTATTTAAA
AAAAGGTGTATTTATAATCAAATACTCGGTACTTNTAATTACTCCAAGAANA
ATTAATTTGAAAAAAAGGGGTTCATTATAAAATATATATTAACCGCTTACAC
20 ATAATCCCCAAACAAAACAGCGATTGGGATTAAAAGGTTCTAAGTCCATCAT
TATAAAAGATCATTTCCGAAAAACAAAAGAAATAGATTCAAAATTAGGCGAC
ATCAGCCCGCTGATAANGATCATAAAAAATACAGAAGCTNATTCAGCGAATCA
GAAANTCCTACTCGCCACTATCCGAAAACNTNGAAAAAAAATGG

25 **Rescue ID** EcoR1
Rescue Sequence 2

TGAAAGGTAGCAACAACGTTTCCTTGGAAAAAGCTGTAAATAGTAAACAAAA
TTGTCAAGTTAACGAGCCAAAGTTATTAAATAAGGTTTCGAGTACGTTGGCATC
GGCTGCCCAGGCAGCAAANAAAAACAAAGACGCAGTTCAAGATCAGCTGGAC
30 ACTTAGAAGANTTTAAGAATTGAAGCACATTNNAAAGAAGANAAACAAGAAC
CCCACCAAAAACCCGCGTGCGTTTGTATGTGTGTGTGCCATCAAATTTCCCGC
ACTGGGTGAATGTGCNTGCGTGTTTGTGTGTCATTTAATTTTCCTACCAATAA
TCGCCTTCCAAGAAGTGAATACCAGCCGATCCACCGCTAAATCGAAAAAAGTT
TNACTCTGGGTAAANTCACTGTTTACGGCTTTTGTGCTATAATTACCTTTCCCG
35 TAAGCNGTGGGAANCTAAAANCCAAAACNTNAGAATCCGAATTCCG

Genomic hit, Accession No. CSC:AC017934

Associated ORF

40 Genscan partial ORF1 predicted sequences
>22:35:21|GENSCAN_predicted_peptide_4|128_aa
MGTN SGATAGINNKPVG GATGAGVLVGGGVGGANSSIGGVLSNSLGGGGSGGLS
ISGLNAGGQNANVGGMGNVGGDDGGNGMVGGGVNNQQATTPQYTIPGILHFIQ
HEWSRFELERSQWDVDRAELQ

45 >22:35:21|GENSCAN_predicted_CDS_4|384_bp

atgggcaccaattcgggagccaccgctggcataaacaacaagccggtggcggtgcaacaggagccggcgctcctttaggcg
gcggtgtgggcggtgccaatcctcgatcggcggtgtcctgtcgaacagcctgggcggtggcggcagcggcggtctgagcatc
agcggcctcaacgctggtggacagaaacgccaatgtgggcggaatgggcaacgttggcggcgacgacggcggaacgggatg
gtgggcggcggtgtaaataaccagcaggccacaacgccccatacacaataccgggcacatctgcacttcacccagcacgagtg
5 tcgcgcttcgagctggagcgatcacagtgggacgtggacagggccgaattgcag

Human Homologue TBLASTN with ORF1: very weak homology with striatin,
calmodulin-binding protein (STRN) (NM_003162.1)
Drosophila EST several including LD42534 (AI516610), LD03224

Annotated Drosophila genome genomic segment AE003619
Annotated Drosophila genome Complete gene candidate CG7392 – novel WD40 family member

Human homologue of Complete gene candidate CG7392- SG2N_HUMAN
CELL-CYCLE NUCLEAR
AUTOANTIGEN SG2NA
(S/G2... 622 e-178 A cell-
cycle nuclear autoantigen
containing WD-40 motifs
expressed mainly in S
and G2 phase cells

Putative function WD40 protein a novel nuclear protein mainly expressed in S and
G2 phase cells that was characterized using autoantibodies from a
cancer patient

Confirmation by RNAi Reduction of G1peak , more polyploidy

Line ID 103/1
Category 2nd chromosome, small imaginal discs
Reversion R
Map Position 57B

Rescue ID BamH1
Rescue Sequence 1

GATTTCAA AATTAGGCGACATCAGCCCGCTGATAAAGAATCATAAAAAATACT
GAGGCTTATTTTAGCGAGTCAGAGACTCCTACTCGCCA ACTATCGAAAACATA
GNGAAGATATAGTCGCCAACCGATCTGCCTTCTATAGTGTTGCTTATTGTTGTC
CCCTAATCAAATTAATAAAAAATCTGCATTAGGCTGCTTCGCCGGCCAGCAACA
AATGTTTTACACCTACTGTACTTTTCGCAGAACAGAGATCCAAATGCAGGATC
GTTTCCATGACTGTACATTTATTCGGATTAGACATTAAATTACACCCTACAGCT
ATACATACTAACAGTGAACACGGCAAATGCTTAGCTAGCATTGGGCCACTTTC
GTTGACTGCGAATAAAAAATGATTGGCCGATGCCTTTAGCAGATTCTTTTGAT
CGAATTACTCGGATGGCTTGTGTGTCCACCTCTTACAAGAACTCCTCGCACCA

ATCGTTGAGACAGTTGTAGCAATCGGATGCTTGGTTGGAGCTGGCGTGGCACA
CCTTCTTCATCCAGTCCTTGGACAGNTTCTTGGNCCTTTTCAGNANCAGGATCT
GGTCCCAAACGGNGGAAGGCCTAAAACGAATGGNAATTGATCGGTAGCCCTT
GACTGGCATTGGTAATTTGCGCACATGGGNGTCATCGGATTTACACACGCACC
5 ATATCGAATCAGCGTCCTTAAGCGTCAACCGAGGGTTTCCCCAATTCCGGCCA
GTTCCGTCACCGACTTGGTTGCCATTGG

Rescue ID **EcoR1****Rescue Sequence 2**

10 ATCAAAGCGNCTGGGCCCCGTGCATCGCCNCAGCGTTCGTCTTAATTAATTAGT
GATTGCAAGCGGGTGCAATTATGCACAAAATTACGGACTAATACAACCTGCCC
GCTTCGCGCTCTCTCCATCTCCCTTCCAAATAGTCGTTTGCTCTTCGCACACAA
AAGTGTA AACCTGTGAAAGGTAGCAACAACGTTTCCTTGGAAAAAGCTGTA
AATAGTAAACAAAATTGTCAAGTTAACGAGCCAAAGTTATTAAATAAGGTTTCG
15 AGTACGTTGGCATCGGCTGCCCAGGCAGCAAAGAAAAACAAAGACGCAGTTC
AAGATTCAGCTGGACACTTAGAAGAGTTTAAGAATTGAAGCACATAAAAAAG
AAGAGAAACAAGAACCCACCAAAAACCCCGCCGTGCGTTTGTATGTGTGTG
TGCCATTCAAATTTCCCTGCACTGGGTGAGTGTGCGTGTGTGTGTGTGTGTC
AGTTTAATTTTCTACCAATAATCGCCTTTCCAAGACGTGATTACCAGCCGATC
20 CACCGCTTAAAATTGATAAACGTTTTTA ACTCTTGCGTTACATCAGCTGTTTTAC
GGCTTTTTGTGCTATAAGTTACGCTTTTCCCGTAAGCCGTTGGCAACACTAGAA
CGCAAAAGAGCATAAAGAATCGCGAGTACCGTANAGAGGAAGAGAGGAAGA
GAGAGAGATAGAGAGTGTGAGCGTGTGAGTGAGCGGGGAATGTGGGGGCGGT
TCCGGTGCGAAAAAACGTAGTAGTAGTACATNNAGAGAGTGCGAACGAGAGG
25 GAGGCAGCCAGCGAGTGTCTGCGACTGCTCCCCCCTTTACCCTCGTCGCTTTT
CTATTCGGAAAATTCAATGACCTCATTTGTTTCATGTGCCGAACCTTTGCTTTTC
TTTCCCAACCTAAAAACGCAAAAAAAAAAAAAACNCCAAACAGGATATACGTNG
GAACANTGANCAAACNANTTCGANAAAACCAACAACNANGGACCGTGCCCTG
GGGCNCCTGAAAGGCAAACAGCTGGCNCNCAAATCCGGAAAAGGATCNGGAA
30 NAACAGGATCNGCGGGCNCAAGGATCNC CGGAACAGGCAAAGGAAACNCCC
GGCNCACNGCACAAGCCNCTGAAAAGCAACNTGAACCAATGGGCACCANTTC
CGGGANCCACCGCTGGCATTAAA

Genomic hit, Accession No. CSC:AC017934

35

rest of results as for line 6/7

Example 66 (Category 5)

Line ID 65/24
Category 2nd chromosome, small imaginal discs
5 Reversion NR
Map Position 48A

Rescue ID BamH1

Rescue Sequence

10 TACGATTTTTGCANTGCNCCATTTCGTGGCACCCGATTTGTATATATATTTTTT
ATATAACCCACGGATTGCCAACTTTCATTGCCCTTTCACACTCTTATTCGCCAT
TTATGAACTCTTCTTTGACGATTGGAACGGTTCTTTTTTCGCTATTTTCGACTGC
ACCCGCGCTCTTTTCGCTTCGCTCTCCTCCCTCTCTACACACCGCTCTTTATCCT
TAATTGCTTTTTTCTATTTAGCGGAATTGATCGTTCTCAACTTGGTCGCCATTGC
15 AGCTCCACAGGCGAAAAAATCGGTGGAAATGCCAATACAGGTGCACGGCGAG
TGCCGATAAGCTGGAAAAATCGGGAAAAACGCACGCCTACACATTCATTGCCAG
CATCGGCTTTGCCTTTTTCGCTGTCGAGATTAGCATATTTCCACTTTTGGTTCGC
GCACAACACTANCTAAATTATTGNTTATTTTTTTTCCCCAACTGTGAGGTGAAAC
TGTGAAACAAAACCACTGTGGGCGGGTCAGTGTGACCCTCTCGCGGTGGGTG
20 AAAATCCTAGTGAGCTTCGTTGTTAGGGCTGTATGACACGAAAGCAAGTTGAA
AAGAACTTTTTTAAAATTATATTGGTTAATTGAGCAGAACTAAACTATATN
AAAATATTTAAGAATNCAGATTAGTGATGTATTTAATATAATAATAGTAAGAT
GTTC

25 Rescue ID EcoR1

Rescue Sequence 2

CTTNTTTGATAGANATAGGCTTCTTTTAAAAAAAANAAGCAGCANCAGGGG
CCCNAGAGTGCGTGNNTGTGAACGCTGATTGCTTGCAAGTGTGTTTCGTGTGTG
TGTGATTGTGTGCTCCGANCAAGTGAAATCAATAATATTTGCAGCCACAAGCA
30 ATTAATAAAAACTGCAATAATGTCAAAAAATCTAATTGAGGCAACAAATTAN
CAAAGCCATNAAAGCAGGCTGCACTGCGAGAAAATTGTGCCTTTCCACAGAT
CTTCTGCTGCAAAGCNAAAGAANGTAAGCAAGTCGGCCANTTTATTNCATTCT
TCTCATCTCTCTTCTTCGCGAATTGGCGCNTANCACTTACAATAATTNATATNA
CTTCTTAAATTTCAAANTCCCTTTCNTGAACGGGANCTTTTAACGGAAAACAAA
35 GCGGGTAAACTAACTTAACTAACTAATTANAANTGTANGTATAAATGAACC
GAACTCGCTTTAGATATNATGCGTTTCACTAACANATTANAACAACTTTGAA
GCTGTANTGTCAGGTTGTTATTNCGTTACCANATGTAGACTGNCCGNNAATT
TNACCTTTCCCATANTCTGTTCTTAANTGTNTTGTTTTTTCCCAATNNTTTGATC
ATNCNTTGGTNAATNANCTNAACGGCCCAAAGTNAATGAATTCCANTCACGTC
40 CACTGGCTCTGGTTCNATANTTAATNGGCTGTTTCTTACTTCCCTTAACCCTAA
CATCTNTTAATCACCTGTGCCATNTGTTTGTGTGTGTGTGAACGAATGAGAAA
AAAAA

Annotated *Drosophila* genome genomic segment AE003825

Annotated *Drosophila* genome Complete gene candidate CG9005 - novel putative cell adhesion

Human homologue of Complete gene candidate CG9005- Ensembl predicted gene
ENSP00000006008
Gene:ENSG00000005238
Clone:AC004472
Contig:AC004472.00001 6.00E-38
(KIAA1539 protein AB040972) and
AK022837 Homo sapiens cDNA
FLJ12775 4e-33

Putative function Putative cell adhesion protein

Confirmation by RNAi Reduced G2/M peak

Example 67 (Category 5)

| | | |
|----|---|---|
| | Line ID | 74/3 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | NR |
| | Map Position | 47A |
| | Rescue ID | EcoR1 |
| | Rescue Sequence | |
| 10 | GCACAGAATGGCNCCTTCACGACAAAAGATCTNCNAATTAGGATGATGCAGA | |
| | AGGAGGACACGCTTTTTCATTATCTGGTTGCCACCTAATTTAAGTTCCACATCAA | |
| | GGGAAGAAGGAAATACGTTCCAACGGACGTCAAATTTACTAACTACACTACTT | |
| | GAAAAGCCTGTCTATAAAAACACGATAACGTTTTTGCTAATCTCAAGACAATG | |
| | TTAAATATAATTGGAGAAAGTATTGAATATGAATATCACAAAAATTGTTTAGG | |
| 15 | GTCTCTACGTGGTAAATAGTATTTGGCATAGACAGTGAGATGTGAGTCGTACG | |
| | TACTAATTAATAAAGTTGTTCAARAGAACCTCATATACTGTAAGTGACAACGA | |
| | ACGAAGCTGACAACCTCTGCTTGACATATTTGGCGGAGTTCGAAAATATCATC | |
| | GCATTGGTATTGTTTTTGTNTCCACCNTGGGGCGAGATTTTGTTGTTGCTTTAC | |
| | TTTGCTTGTTTTTTCNCCACAAANCGAACCATAATGTTTCGAAATGGTAAAATTA | |
| 20 | CCGTGCCAACAAGCTCTCTCTCTCCCCACTCCGAAACTCTCTCATCTCTCCTTG | |
| | CAATTGTTTAAGGTGTGCAAGGAAATGAAAAATGTCCCGGCTGTGTTNCCATG | |
| | CATTCCCCTTCAAAGCCAATTATNTTTGTGCCTCTCCAACNTTTTTGATCGGNN | |
| | TGATTTTTTTGGCTCCCCNTANTCCCCCCCCCTTTCNCCCATTCCGGGTTANAT | |
| | TATTNTNCCAATTTTCCTATTTTACGGTCCCNGTTCCCTGGAAATANTTCCTNC | |
| 25 | AATCNCCGCTCCATNTCNCCATNTTTGACAGATTTTC | |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003829 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG12052 lola -a specific RNA |
| 30 | | polymerase II transcription |
| | | factor involved in axon |
| | | guidance |
| | Human homologue of Complete gene candidate | 1e-09 3789797 (AF059569) |
| 35 | | actin binding protein |
| | | MAYVEN [Homo sapiens] |
| | Putative function | lola-like specific RNA polymerase II transcription factor |
| | Confirmation by RNAi | Almost no G1 peak and increase in G2/M peak indicating |
| 40 | | arrest in G2/M |

Example 68 (Category 5)

| | | |
|----|--|--------------------------------------|
| | Line ID | 79/7 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | R |
| | Map Position | 55B |
| | Rescue ID | BamH1 |
| | Rescue Sequence 1 | |
| 10 | GTCTCATGCACCCTGGCCCTNAGCTGCATAAGTGTAAGTGTGTGNCTGTGTGC GAGTGTGGGTAGGCGGCGGCAACTATCTCGCTTGCTCTTGCGTCCGGGGTTAT CGGTAGCTTCTTCTAGGCTGAGTGCATTTCGTTGAATCGTGGATGTTGAAAGTT GTCTAATTTCCGAACCTATTGATTTTTTCCCCTTCCCCGTCAAGAACTGCATTGT TGCTTCTTGAAGACCAGTTTTTGGTAACATCAGGAGAATGGAAAGGAGCGAGT | |
| 15 | GAGTCGGTGAGTAAGTGAGTGAGCGATGCGAGCGACAAAATCAACAACAACA ACAACAACGGTCAAAACGAGTTCCAACGAAAGTTGCAACACTCTCAACAATT TGAGCAGCTCCGTTTGTGTTATTGCATTACTCAATCGGGAAGACTCTACACTC GACGGAATAGTGTGCTCGTCTGAAATTTATCNATTTCCATTCCCTTCCTTTGTTT TTGGGCCAAACAATGGCNTCGGCAANCGTTCGTGGAAAACCGCAGGAACCAC | |
| 20 | CAAAATGCCTGGCGTCACATTAACCGAGCCGCCTTTGTTTATGCAAATATTATT GTAATATTTGGTNAAAATTAAGTCGCGCTTCNCGTTACTTTTTATTTCATATAC ACGCAGCAGCAGCACGCATACAGTCACGTCACGCACACATACAATCGCCGTN CACATACACTTGTCTTTTTTNCCACACACTTTCCTAATCAT | |
| 25 | Rescue ID | EcoR1 |
| | Rescue Sequence 2 | |
| | NGNGTCTCATGCACCCTGGCCCTNAGCTGCATAAGTGTAAGTGTGTGNCTGT GTGCGAGTGTGGGTAGGCGGCGGCAACTATCTCGCTTGCTCTTGCGTCCGGGG TTATCGGTAGCTTCTTCTAGGCTGAGTGCATTTCGTTGAATCGTGGATGTTGAA | |
| 30 | AGTTGTCTAATTTCCGAACCTATTGATTTTTTCCCCTTCCCCGTCAAGAACTGCA TTGTTGCTTCTTGAAGACCAGTTTTTGGTAACATCAGGAGAATGGAAAGGAGCG AGTGAGTCGGTGAGTAAGTGAGTGAGCGATGCGAGCGACAAAATCAACAACA ACAACAACAACGGTTCAAAACGAGTTCCAACGAAAGTTGCAACACTCTCAAC AATTTGAGCAGCTCCGTTTGTGTTATTGCATTACTCAATCGGGAAGAACTCTA | |
| 35 | CACTCGACGGAATAGTGTGCTCGTCTGAAATTTATCNATTTCCATTCCCTTCCTT TGTTTTTGGGCCAAACAATGGCNTCGGCAANCGTTCGTGGAAAACCGCAGGA ACCACCAAATGCCTGGCGTCACATTAACCGAGCCGCCTTTGTTTATGCAAAT ATTATTGTAATATTTGGTNAAAATTAAGTCGCGCTTCNCGTTACTTTTTATTTC ATATACACGCAGCAGCACGCATACAGTCACGTCACGCACACATACAATCGCC | |
| 40 | GTNCACATACACTTGTCTTTTTTNCCACACACTTTCCTAATCATNNTA | |
| | Genomic hit, Accession No. AC004296 | |
| | Associated ORF | |
| 45 | Genscan: ORF2 predicted sequences >15:31:31 GENSCAN_predicted_peptide_3 109_aa MVTSEFRHLRDEKSFTDVTLACEGQTCKAHKMVLSACSPYFKALLEENPSKHPIIL | |

KDVSYIHLQAILEFMYAGEVNVSQEQLP AFLKTADRLKVKGLAETPSSIKREG

>15:31:31|GENSCAN_predicted_CDS_3|330_bp
atggtgacctcggtccgtcacctgcgcgacgagaagagcttcacagatgtaacactcgctgcgagggccaaacctgcaaagcc
5 caaaaatggtgcttccgcttgacgtccctactttaagcgctactggaggagaacccatcgaagcatccgatcattatcctgaaa
gatgtctctacattcacctacaggctatactggagttcatgtacgccggtgaggtgaacgtgtcccaggaacaattgccagcatt
cttaagaccgccgatcgctcaaagtgaaggcctcgacagacacccagttcgataaagcggggaaggtga

Drosophila Gene Hit TBLASTN with ORF2: several zinc finger proteins including
10 Broad-Complex mRNA for BRcore-Z2 protein (X54665)
Human Homologue TBLASTN with ORF2: kelch (*Drosophila*)-like 2 (Mayven actin
binding protein) (KLHL2) (AF059569)

Annotated Drosophila genome genomic segment AE003800
15 **Annotated Drosophila genome Complete gene candidate** CG5738- lolal, lola-like
putative kelch-like putative
specific RNA polymerase II
transcription factor known to
affect disc morphology
20
or could be CG10914 - novel
unknown

Human homologue of Complete gene candidate CG5738- 9e-09 3789797
25 (AF059569) actin binding
protein MAYVEN [Homo
sapiens]

CG10914- predicted gene
30 ENSP00000051207
Gene:ENSG00000047313
Clone:AC068261
Contig:AC068261.00019
4.00E-49 (potential cell
35 division GTP binding protein
1: ENST00000051207

Putative function CG5738: lola like specific RNA polymersae II transcription factor,
40 CG10914: Possible GTP binding protein

Confirmation by RNAi Both show marked reduction in G1 to G2/M ratio

Example 69 (Category 5)

| | | |
|----|---|--------------------------------------|
| | Line ID | 80/2, 81/8 |
| | Category | 2nd chromosome, small imaginal discs |
| 5 | Reversion | R |
| | Map Position | 57D/E |
| | Rescue ID | BamH1 |
| | Rescue Sequence 1 | |
| 10 | CANTTTCAGAGGCCATAGNCCTTCACAAAATTCNCCATCTCTGCCCCGGCATCC GTGCTTGAAAATGGTGCCAATGCGTCGTGGAGAATCTGCTGCACTCGATGGTC TGCAAAATTGCACATTTATTAGATTTAATAAATTTTTCAACTGTCCGCGANCAC GTTTGCTCGTGTTGAATTTTCGAGTACAAAATTAGTGCGACTGTTGGATTGCATT GAAATGCCAAAAATCGGTGTGACCATTTCGAAGTCCCCACAGGCTCATGACTT | |
| 15 | TCGCGGTTACCAAATCCAAATAACGCAAGCTGGTCACGCTGTCAAACATCGG TGACGGAATGGTGACGACACAAACAATTTGCTTAAAAAATTTCTTGCGGCCGT AAAAATGCGCAAGCAGCCTGGCAGCGCAACGCACGTACACGTAATTGGAACA AATGTTTGCTGAACCACAACCGCCCACTAAATGTTANCCGCCAAGTCTTTTCC CCCGCCGCCGCCGTCNTCNCNCNCCGGATTATTTGGTTTACAATTTGCTTAC | |
| 20 | ACAAGTGCAATCGTCGATAGCGCTTCATTTTGGAGTAACAAGTAATATTTTGC GCCGTACTGCTGTTTCGCCGTATCAGACAGAAGGTTGGTATCAGTTCGACGCAG CTTGTGACGGTATTGCATACGCGGGCGAAACGCCACGTGAAAACGGATCGCA GTTCTCGAAAACTCNGGATAAAAAA | |
| 25 | Rescue ID | EcoR1 |
| | Rescue Sequence 2 | |
| | TGGGGTCTCANGCCCCGACGGCCATATTTTAAACACAAGATTCNNCANCTCTGC AGGGCATCCGTGCTTGAAAATGGTGCCAATGCGTCGTGGAGAATCTGCTGCAC TCGATGGTCTGCAAAATTGCACATTTATTAGATTTAATAAATTTTTCAACTGTC | |
| 30 | CGCGAGCACGTTTGCTCGGTGTTGAATTTTCGAGTACAAAATTAGTGCGACTGT TGGATTGCATTGAAATGCCAAAAATCGGTGTGACCATTTCGAAGTCCCCACAG GCTCATGACTTTCGCGGTTACCAAATCCAAATAACGCAAGCTGGTCACGCTG TCAAACATCGGTGACGGAATGGTGACGACACAAACAATTTGCTTAAAAAATTT CTTGCGGCCGTAAAAATGCGCAAGCAGCCTGGCAGCGCAACGCACGTACACG | |
| 35 | TAATTGGAACAAATGTTTGCTGAACCACAACCGCCCACTAAATGTTAGCGCCA ACTNCTTTTCCCCGCCGCCGGTCGTCNTCNCNCCGGATTATTTTGTGTTACA ATTTGCTTACACAAGTGCAATCGTCGATAGCGCTTCATTTTGGAGTAACAAGT AGTATTTTGCGCCGTACTGCTGTTTCGCCGTATCANACAGAAGGTTGGTATCAG TTCGACGCAGCTTGTGACGGTATGCATACGCGGGGAAACGCCACGTGAAAAC | |
| 40 | GGATCGCAGTNCTCGAAACTCNGGATAAAAGAAAAAGTAGGCTGAATG | |
| | Genomic hit, Accession No. AC007175 | |
| | Associated ORF | |
| 45 | Genscan: ORF2 predicted sequences >16:09:09 GENSCAN_predicted_peptide_3 2497_aa MNEGNSAGGGHEGLSPAPPAVPDRVTPHSTEISVAPANSTSTTVRAAGSVGAALP | |

ATRHHQHIAATQVKGIASSSSKQQKQLASAPVPLSPLPQQQQQTAEATAAAAAP
AHSNVSVSSSTIEASVLPQAKRQRLDDNEDRTSAASIVGPAESSNIVSSLLPASVA
SSSEVGGLSSTALQDLNALKKRILQQKLQILRNLKERHLENVSEYFYLQNGGSM
DYPAWRKKTPTPQFISYSNANRIDQLIHEDKPSTSAAAAAAQNQKYTTQQTDSVE
5 SSLVSGIGTGATKGAPLDGNISNSTVKTNTQSQVPSKIGSFTESTPAATESNSSTTV
GTATSGAATSTSATSASGNVLAVEAEIKIPAVGATPVAISTKLPAAVVQLTQQG
GTPLPCNTSAGSTALRRPQGQNNASSGSAAASGGGGSLTPTPLYTGNGPAALGG
SGGLTPGTPTSGSLLSPALGGGSGTPNSAAQEFSFKAKQEVYVMQRISLQREGL
WTERRLPKLQEPSRPAHWDYLLEEMVWLAADFAQERKWKKNAAKKCAKMOV
10 QKYFQDKATAAQRAEKAQELQLKRVASFIAREVKSFWSNVEKLVEYKHQTKIEE
KRKQALDQHLSFIVDQTEKFSQQLEGMNKSVAADTPSLNSSRLTSPKRESDDDFR
PESGSEDDEETIAKAEEDAADVKEEVTALAKESEMDFDLNDLPPGYLENRDKL
MKEEQSSAIKTETPDDSDSEFEAKEASDDDDENTISKQEEAEQEIDHKKEIDELEA
DNDLSVEQLLAKYKSEQPPSPKRRKLAPRDPELDSDDDSTAVDSTEESDAATED
15 EEDLSTVKTDMDMEEQDEQEDGLKSLMADADATSGAAGSGSTAGASGNKDDML
NDAAALAESLQPKGNTLSSTNVVTPVPFLLKHSREYQHIGLDWLVTMNERKLN
GILADEMGLGKTIQTIALLAHLACAKGNWGPFLVVPSSVMLNWEMEFKKWCPG
FKILTYYGSKERKLKRVGWTKPNAFHVCSYKLVVQDQQSFRKKWKYLILD
EAQNIKNFKSQRWQLLLNFSTERRLLLTGTPLQNDLMELWSLMHFLMPYVFSSHR
20 EFKEWFSNPMTGMEGNMEYNETLITRLHKVIRPFLRLKKEVEKQMPKKYEHV
ITCRLSNRQRYLYEDFMSRAKTRETLQTGNLLSVINVLMQLRKVCNHPNMFEARP
TISPFQMDGITFHTPRLVCDIMEYDPFTQINLETNLLHLEQTMAYVSHKSRLL
APPRKLIEDIDTAPLPAPRCPNGKYRFHIRVRSALAQRILNAVKGASPAAMRLE
GSKIMPMRNLLPSGRVLKRVASINPVNMALKPVVINSVVTTTSSSTTASSPTGAL
25 SVLSNSKLLGARSQINAPTPAKVAKTMQDGKPFYLTATNSGAAGARLTLSKT
TASASTTTSRTTVTASTTSGQQLIRDPIVKDLATHVKSTVQKQSIANGKTEPEEETE
AEDPYKVQELIQMRKEQRLAALKRMAMINRRRTDATPIYGEDCREAIQRCMQAT
RSLKRSTWQTRGYANCCTAMAHNRNGWSLNHLLKSFEERCADLKPVFANFVIYVP
SVCAPRIRRYVQNLSSTHWQHEQRIENIVDQALRPKLALLHPIISEMTTKFPDPRLI
30 QYDCGKLQTMDRLLRQLKVNGHRVLIFTQMTKMLDVLEAFLNYHGHYLRDLGS
TRVEQRQILMERFNGDKRIFCFILSTRSGGVGINLTGADTVIFYDSWNPTMDAQA
QDRCHRIGQTRDVHIYRLVSERTIEVNILKKANQKRMLSDMAIEGGNFTTTYFKSS
TIKDLFTMEQSEQDESSQEKSENKDRIVATTTLSSTPSTVVETEKQSLRAFEHALA
AAEDEQDVQATKTAKAEVAADLAEFDENIPIATEDPNAEGGPQVELSKADLEMQ
35 NLVKQLSPIERYAMRFVEETGAAWTAEQLRAAEAELEAQKREWEANRLAAMHK
EEELLKQETEAEEMLTYSRKDSSNQVNTKTDSSNSNKRRLVRENRRNSAQKLSRSV
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 40 gtgggtgttcgggagggaatgcctcctcgagcgggaacagccaggtga

Drosophila Gene Hit TBLASTN with ORF2: brahma protein (M85049) and imitation-SWI protein (ISWI) (L27127) and chromodomain-helicase-DNA-binding (CHD-1)

45 **Human Homologue** BLASTX with EST TBLASTN with ORF2: Snf2-related CBP activator protein (SRCAP) (AF143946) and SWI/SNF related, matrix associated, actin dependent regulator of chromatin, subfamily a, member 4 (SMARCA4) (NM_003072.1)

| | | |
|----|---|---|
| | <i>Drosophila</i> EST | several including SD07794 (AI534784), LD34465 (AA990657) |
| | Annotated <i>Drosophila</i> genome genomic segment | AE003453 |
| 5 | Annotated <i>Drosophila</i> genome Complete gene candidate | CG9696 – domino an enzyme involved in DNA repair homology to snf2 family helicases |
| 10 | Human homologue of Complete gene candidate | CG9696- gi4557447 416409C913D6A935 ref NP_001261.1 chromodomain helicase DNA binding protein 1 [Homo sapiens] (1.90E-85 |
| 15 | Putative function | snf2 helicase family member protein that contains a chromodomain, which occurs in proteins that are implicated in chromatin compaction, and an SNF2/SWI2-like helicase domain, which occurs in proteins that are believed to activate transcription by counteracting the repressive effects of chromatin structure |
| 20 | | |
| | Confirmation by RNAi | Loss of G1, peak, increae in G2M indicating arrest in G2/M |
| 25 | | |

Example 70 (Category 5)

Line ID 99/31
Category 2nd chromosome, small imaginal discs
5 **Reversion** NR
Map Position 53E

Rescue ID EcoR1
Rescue Sequence 1

10 AAGGCCCGACCAGAAACGAAATTTTCGGCGCGTNTTTTAAAATGCGCGGTAA
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25 ACATTCCNGGCTTTCCCAATTTTCNCCTTTACTACAATTTCAATGGTTTCTTTTT
CCTCAC

Rescue ID BamH1
Rescue Sequence 2

30 CCTNAAATGTNGCGCTGGGNCCTAAANCGTCNCTCCTTGTGTCTCTCTTGTTTA
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40 TGAAGAGAGCGAGAAGAAGAAANCGANATGAGATGCAGAGGACAGATAAAG
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Genomic hit, Accession No. CSC:AC020063
45 **Associated ORF**

Genscan ORF1 predicted sequences >16:48:25|GENSCAN_predicted_peptide_1|722_aa
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 5 DESKPKSGADKPKKPEPKAKNGKVAKEEDDDDEDEDEDEDAEDDDGDENDGLDK
 NNEVAEDDENVVVALAEIDRINENINKTRVDGLQTLHAICFGAQGKNNVVKKNLRS
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 EPDESLCLEQGDEEEEEEDAEDEDLDEDEEDPPSEEDKKRKSGKSSGGAGRGSARN
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 10 SDGGRGGGAGAAGRKVPSRGGRRPARKSRRRNSDSEEEEESEVSDADSDVPKR
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 15 IWLICCCNNQIFGET

>16:48:25|GENSCAN_predicted_CDS_1|2169_bp
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 20 ccccgccgctgtcgacgcctctgcctctgccgccactgatgacgtcgctgataaaaaagccaaaggagactcaacggctgttca
 aataccgaatcgatgcagctgcagcggacaaaaaggagaaatccccctgccggtaataaagaagtccaacaataaggatgc
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45 **Human Homologue** TBLASTN with ORF1: poor homology with DEK gene
 (D6S231E) (NM_003472.1)
Drosophila EST several including LD33301 (AA979048)

| | | |
|----|---|---|
| | Annotated <i>Drosophila</i> genome genomic segment | AE003805 |
| | Annotated <i>Drosophila</i> genome Complete gene candidate | CG5935 - EG:EG0003.6 - novel with weak homology to DEK oncogene |
| 5 | | CG8648 - EG:EG0003.3 - novel XPG/ flap endonuclease-like, DNA repair? |
| 10 | Human homologue of Complete gene candidate | CG5935- 1e-17 4503249 ref NP_003463.1 pD6S231E DEK gene >gi 544150 sp P35659 DEK_H UMAN DEK PROTEIN >gi 284375 |
| 15 | | CG8648- 4758356 ref NP_004102.1 pFEN1 flap structure-specific endonuclease 1; MATURATION FACTOR 1 (MF1); DNase IV; RAD2_HUMAN(aa) |
| 25 | Putative function | CG5935: function unknown but putative DNA-binding protein predicted to be involved in chromosomal organisation. The translocation (6;9), associated with a specific subtype of acute myeloid leukemia, results in the fusion of two genes, dek and can, and the expression of a chimeric, leukemia-specific dek-can mRNA |
| 30 | | CG8648: Novel XPG/ flap endonuclease-like, DNA repair protein |
| | Confirmation by RNAi | Both show slight reduction of G1 peak |

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Each of the applications and patents mentioned above, and each document cited or referenced in each of the foregoing applications and patents, including during the
5 prosecution of each of the foregoing applications and patents (“application cited documents”) and any manufacturer’s instructions or catalogues for any products cited or mentioned in each of the foregoing applications and patents and in any of the application cited documents, are hereby incorporated herein by reference. Furthermore, all documents cited in this text, and all documents cited or referenced in documents cited in this text, and
10 any manufacturer’s instructions or catalogues for any products cited or mentioned in this text, are hereby incorporated herein by reference.

Various modifications and variations of the described methods and system of the invention will be apparent to those skilled in the art without departing from the scope and spirit of the invention. Although the invention has been described in connection with
15 specific preferred embodiments, it should be understood that the invention as claimed should not be unduly limited to such specific embodiments. Indeed, various modifications of the described modes for carrying out the invention which are obvious to those skilled in molecular biology or related fields are intended to be within the scope of the following claims.

CLAIMS

1. A polynucleotide selected from:
 - (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 1 to 70 or the complement thereof.
 - 5 (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 1 to 70, or a fragment thereof.
 - (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 1 to 70 or a fragment thereof.
 - 10 (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).
2. A polynucleotide selected from:
 - (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 1 to 14 or the complement thereof.
 - 15 (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 1 to 14, or a fragment thereof.
 - (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 1 to 14 or a fragment thereof.
 - 20 (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).
3. A polynucleotide selected from:

- (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 15 to 19 or the complement thereof.
- (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 15 to 19, or a fragment thereof.
- 5 (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 15 to 19 or a fragment thereof.
- (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).
- 10 4. A polynucleotide selected from:
- (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 20 to 30 or the complement thereof.
- (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 20 to 30, or a fragment thereof.
- 15 (c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in Examples 20 to 30 or a fragment thereof.
- (d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).
- 20 5. A polynucleotide selected from:
- (a) polynucleotides comprising any one of the nucleotide sequences set out in Examples 31 to 53 or the complement thereof.
- (b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in Examples 31 to 53, or a fragment thereof.

(c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in 31 to 53 or a fragment thereof.

(d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

6. A polynucleotide selected from:

(a) polynucleotides comprising any one of the nucleotide sequences set out in 54 to 70 or the complement thereof.

(b) polynucleotides comprising a nucleotide sequence capable of hybridising to the nucleotide sequences set out in 54 to 70, or a fragment thereof.

(c) polynucleotides comprising a nucleotide sequence capable of hybridising to the complement of the nucleotide sequences set out in 54 to 70 or a fragment thereof.

(d) polynucleotides comprising a polynucleotide sequence which is degenerate as a result of the genetic code to the polynucleotides defined in (a), (b) or (c).

7. A polynucleotide probe which comprises a fragment of at least 15 nucleotides of a polynucleotide according to any of Claims 1 to 6.

8. A polypeptide which comprises any one of the amino acid sequences set out in Examples 1 to 70 or in any of Examples 1 to 14, Examples 15 to 19, Examples 20 to 30, Examples 31 to 53 and Examples 54 to 70, or a homologue, variant, derivative or fragment thereof.

9. A polynucleotide encoding a polypeptide according to Claim 8.

10. A vector comprising a polynucleotide according to any of Claims 1 to 7 and 9.

11. An expression vector comprising a polynucleotide according to any of Claims 1 to 7 and 9 operably linked to a regulatory sequence capable of directing expression of said polynucleotide in a host cell.

12. An antibody capable of binding a polypeptide according to Claim 8.

5 13. A method for detecting the presence or absence of a polynucleotide according to any of Claims 1 to 7 and 9 in a biological sample which comprises:

(a) bringing the biological sample containing DNA or RNA into contact with a probe according to Claim 9 under hybridising conditions; and

10 (b) detecting any duplex formed between the probe and nucleic acid in the sample.

14. A method for detecting a polypeptide according to Claim 8 present in a biological sample which comprises:

(a) providing an antibody according to Claim 12;

15 (b) incubating a biological sample with said antibody under conditions which allow for the formation of an antibody-antigen complex; and

(c) determining whether antibody-antigen complex comprising said antibody is formed.

15. A polynucleotide according to according to any of Claims 1 to 7 and 9 for use in therapy.

20 16. A polypeptide according to Claim 8 for use in therapy.

17. An antibody according to Claim 12 for use in therapy.

18. A method of treating a tumour or a patient suffering from a proliferative disease comprising administering to a patient in need of treatment an effective amount of a polynucleotide according to any of Claims 1 to 7 and 9.

19. A method of treating a tumour or a patient suffering from a proliferative disease,
5 comprising administering to a patient in need of treatment an effective amount of a polypeptide according to Claim 8.

20. A method of treating a tumour or a patient suffering from a proliferative disease, comprising administering to a patient in need of treatment an effective amount of an antibody according to Claim 12 to a patient.

10 21. Use of a polypeptide according to Claim 8 in a method of identifying a substance capable of affecting the function of the corresponding gene.

22. Use of a polypeptide according to Claim 8 in an assay for identifying a substance capable of inhibiting the cell division cycle.

15 23. Use as claimed in Claim 22, in which the substance is capable of inhibiting mitosis and/or meiosis.

24. A method for identifying a substance capable of binding to a polypeptide according to Claim 8, which method comprises incubating the polypeptide with a candidate substance under suitable conditions and determining whether the substance binds to the polypeptide.

20 25. A method for identifying a substance capable of modulating the function of a polypeptide according to Claim 8 or a polypeptide encoded by a polynucleotide according to any of Claims 1 to 7 and 9, the method comprising the steps of: incubating the polypeptide with a candidate substance and determining whether activity of the polypeptide is thereby modulated.

26. A substance identified by a method or assay according to any of Claims 21 to 25.
27. Use of a substance according to Claim 26 in a method of inhibiting the function of a polypeptide.
28. Use of a substance according to Claim 26 in a method of regulating a cell division
5 cycle function.